

STIC Search Report Biotech-Chem Library

STIC Database Tracking Number

TO: Joseph F Murphy

Location: REM/4D78/4C70

Art Unit: 1646

Thursday, August 18, 2005

Case Serial Number: 09/765534

From: Edward Hart

Location: Biotech-Chem Library

REM-1A55

Phone: 571-272-2512

edward.hart@uspto.gov

Search Notes

Examiner Murphy,

Here are the results of the search you requested.

Please feel free to contact me if you have any questions.

Edward Hart



STIC-Biotech/ChemLib

162513

From:

Murphy, Joseph

· Sent:

Monday, August 15, 2005 11:07 AM STIC-Biotech/ChemLib

To:

Subject:

09765534

STIC/Biotech:

Please do an oligo search of SEQ ID NO: 2, 4.

Please include an interference search.

Please send the results on DISK.

Thanks a lot...

Joseph F. Murphy, Ph.D. Primary Examiner, Art Unit 1646 joseph.murphy@uspto.gov Remsen 4D78 Mailbox: 4C70 (571) 272-0877

STAFF USE ONLY

Searcher: Searcher Phone: 2-

Date Searcher Picked up Date Completed:_

Searcher Prep/Rev. Time: Online Time:_

Type of Search

NA#:_ AA#: SPDI: Interference: Oligomer: S/L:_ Encode/Transl: Structure#:_ Text: Inventor:____ Litigation:_

******* Vendors and cost where applicable STN:

DIALOG: QUESTEL/ORBIT:

LEXIS/NEXIS: SEQUENCE SYSTEM:

WWW/Internet: Other(Specify):_

09765534 Resultsoligo

SEQ ID NO: 2

SUMMARIES

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3	6857	100.0	1298	2	ADP90836	Adp90836 Human fms
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5	6857	100.0	1298	3	AAY70746	Aay70746 Human tyr
6	6857	100.0	1298	5	ABG32043	Abg32043 Human Flt
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9	6857	100.0	1298	6	ABU04296	Abu04296 Human exp
10	6857	100.0	1298	6	ABU04274	Abu04274 Human exp
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15	6857	100.0	1304	6	ABU04279	Abu04279 Human exp
16	6852	99.9	1363	2	ADD90838	-
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25	6852	99.9	1363	6	ABU04295	Abu04295 Human exp
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Issued Patents:

SUMMARIES

Result No.	Score	Query Match	Length I	DB	ID	Description
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2	6857	100.0	1298	1	US-08-340-011-2	Sequence 2, Appli
3	6857	100.0	1298	3	US-08-901-710-2	Sequence 2, Appli
4	6857	100.0	1298	3	US-08-446-648-33-	Sequence 33, Appl
5	6857	100.0	1298	4	US-09-982-610-33 -	Sequence 33. Appl

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RESULT 1
US-08-222-616-33
; Sequence 33, Application US/08222616
; Patent No. 5635177
; GENERAL INFORMATION:
     APPLICANT: Bennett, Brian D.
     APPLICANT: Goeddel, David
    APPLICANT: Lee, James M.
APPLICANT: Matthews, William
APPLICANT: Tsai, Siao Ping
     APPLICANT: Wood, William I.
     TITLE OF INVENTION: PROTEIN TYROSINE KINASE AGONIST TITLE OF INVENTION: ANTIBODIES
    NUMBER OF SEQUENCES: 42
     CORRESPONDENCE ADDRESS:
      ADDRESSEE: Genentech, Inc.
      STREET: 460 Point San Bruno Blvd
      CITY: South San Francisco
      STATE: California
       COUNTRY: USA
      ZIP: 94080
     COMPUTER READABLE FORM:
      MEDIUM TYPE: 5.25 inch, 360 Kb floppy disk
       COMPUTER: IBM PC compatible
      OPERATING SYSTEM: PC-DOS/MS-DOS
       SOFTWARE: patin (Genentech)
     CURRENT APPLICATION DATA:
      APPLICATION NUMBER: US/08/222,616
       FILING DATE: 4-APR-1994
       CLASSIFICATION: 530
     PRIOR APPLICATION DATA:
     APPLICATION NUMBER: PCT/US93/00586
      FILING DATE: 22-JAN-1993
     PRIOR APPLICATION DATA:
     APPLICATION NUMBER: 07/826935
      FILING DATE: 22-JAN-1992
    ATTORNEY/AGENT INFORMATION:
      NAME: Lee, Wendy M.
      REGISTRATION NUMBER:
      REFERENCE/DOCKET NUMBER: 821P2
     TELECOMMUNICATION INFORMATION:
       TELEPHONE: 415/225-1994
       TELEFAX: 415/952-9881
  TELEX: 910/371-7168
INFORMATION FOR SEQ ID NO: 33:
    SEQUENCE CHARACTERISTICS:
       LENGTH: 1298 amino acids
       TYPE: amino acid
       TOPOLOGY: linear
US-08-222-616-33
 Query Match 100.0%; Score 6857; DB 1; Length 1298; Best Local Similarity 100.0%; Pred. No. 0;
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0; Indels

0; Gaps

Matches 1298; Conservative 0; Mismatches

1298 4 US-09-169-079-2

Sequence 2, Appli

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DР			
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Qу	241	ELLVGEKLVLNCTVWAEFNSGVTFDWDYPGKQAERGKWVPERRSQQTHTELSSILTIHNV	300
Db	241		300
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Db		HDKHCHKKYLSVQALEAPRLTQNLTDLLVNVSDSLEMQCLVAGAHAPSIVWYKDERLLEE	
ДУ		KSGVDLADSNQKLSIQRVREEDAGRYLCSVCNAKGCVNSSASVAVEGSEDKGSMEIVILV	, , ,
Db		KSGVDLADSNQKLSIQRVREEDAGRYLCSVCNAKGCVNSSASVAVEGSEDKGSMEIVILV	
Qy Db		GTGVIAVFFWVLLLLIFCNMRRPAHADIKTGYLSIIMDPGEVPLEEQCEYLSYDASQWEF	
Qy		PRERLHLGRVLGYGAFGKVVEASAFGIHKGSSCDTVAVKMLKEGATASEHRALMSELKIL	
Db			
Qy		IHIGNHLNVVNLLGACTKPQGPLMVIVEFCKYGNLSNFLRAKRDAFSPCAEKSPEQRGRF	
Db			
Qу	961	RAMVELARLDRRRPGSSDRVLFARFSKTEGGARRASPDQEAEDLWLSPLTMEDLVCYSFQ	1020
Db	961		1020
Qу	1021	VARGMEFLASRKCIHRDLAARNILLSESDVVKICDFGLARDIYKDPDYVRKGSARLPLKW	1080
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RESULT 4
US-08-446-648-33
; Sequence 33, Application US/08446648
; Patent No. 6331302
  GENERAL INFORMATION:
    APPLICANT: Genentech, Inc.
    APPLICANT: Bennett, Brian D.
    APPLICANT: Goeddel, David
    APPLICANT: Lee, James M.
    APPLICANT: Matthews, William
    APPLICANT: Tsai, Siao Ping
    APPLICANT: Wood, William I.
    TITLE OF INVENTION: PROTEIN TYROSINE KINASE AGONIST ANTIBODIES
    NUMBER OF SEQUENCES: 45
    CORRESPONDENCE ADDRESS:
      ADDRESSEE: Genentech, Inc.
      STREET: 460 Point San Bruno Blvd
      CITY: South San Francisco
      STATE: California
      COUNTRY: USA
      ZIP: 94080
    COMPUTER READABLE FORM:
      MEDIUM TYPE: 3.5 inch, 1.44 Mb floppy disk
      COMPUTER: IBM PC compatible OPERATING SYSTEM: PC-DOS/MS-DOS
      SOFTWARE: WinPatin (Genentech)
    CURRENT APPLICATION DATA:
      APPLICATION NUMBER: US/08/446,648
      FILING DATE:
      CLASSIFICATION: 435
    PRIOR APPLICATION DATA:
      APPLICATION NUMBER: 08/222616
      FILING DATE: 04-APR-1994
    ATTORNEY/AGENT INFORMATION:
      NAME: Lee, Wendy M.
      REGISTRATION NUMBER: 40,378
      REFERENCE/DOCKET NUMBER: P0821P3PCT
    TELECOMMUNICATION INFORMATION:
      TELEPHONE: 415/225-1994
      TELEFAX: 415/952-9881
      TELEX: 910/371-7168
  INFORMATION FOR SEQ ID NO: 33:
    SEQUENCE CHARACTERISTICS:
      LENGTH: 1298 amino acids
      TYPE: Amino Acid
      TOPOLOGY: Linear
US-08-446-648-33
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; Sequence 33, Application US/09982610
; Patent No. 6673343
   GENERAL INFORMATION:
         APPLICANT: Genentech, Inc.
                    Bennett, Brian D.
                    Goeddel, David
                     Lee, James M.
                     Matthews, William
                     Tsai, Siao Ping
                     Wood, William I.
         TITLE OF INVENTION: PROTEIN TYROSINE KINASE AGONIST ANTIBODIES
         NUMBER OF SEQUENCES: 45
         CORRESPONDENCE ADDRESS:
              ADDRESSEE: Genentech, Inc.
              STREET: 460 Point San Bruno Blvd
              CITY: South San Francisco
              STATE: California
              COUNTRY: USA
              ZIP: 94080
         COMPUTER READABLE FORM:
              MEDIUM TYPE: 3.5 inch, 1.44 Mb floppy disk
              COMPUTER: IBM PC compatible
              OPERATING SYSTEM: PC-DOS/MS-DOS
              SOFTWARE: WinPatin (Genentech)
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              APPLICATION NUMBER: US/09/982,610
              FILING DATE: 17-Oct-2001
              CLASSIFICATION: <Unknown>
         PRIOR APPLICATION DATA:
              APPLICATION NUMBER: 08/446,648
              FILING DATE: 1996-MAY-23
              APPLICATION NUMBER: 08/222616
              FILING DATE: 04-APR-1994
         ATTORNEY/AGENT INFORMATION:
              NAME: Lee, Wendy M.
              REGISTRATION NUMBER: 40,378
              REFERENCE/DOCKET NUMBER: P0821P3PCT
         TELECOMMUNICATION INFORMATION:
              TELEPHONE: 415/225-1994
              TELEFAX: 415/952-9881
              TELEX: 910/371-7168
    INFORMATION FOR SEQ ID NO: 33:
         SEQUENCE CHARACTERISTICS:
              LENGTH: 1298 amino acids
              TYPE: Amino Acid
              TOPOLOGY: Linear
         SEQUENCE DESCRIPTION: SEQ ID NO: 33:
US-09-982-610-33
  Query Match 100.0%; Score 6857; DB 4; Length 1298; Best Local Similarity 100.0%; Pred. No. 0;
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; Sequence 33, Application PC/TUS9504228
  GENERAL INFORMATION:
     APPLICANT: Genentech, Inc.
APPLICANT: Bennett, Brian D.
     APPLICANT: Goeddel, David
     APPLICANT: Lee, James M. APPLICANT: Matthews, William
     APPLICANT: Tsai, Siao Ping
     APPLICANT: Wood, William I.
     TITLE OF INVENTION: PROTEIN TYROSINE KINASE AGONIST ANTIBODIES
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NUMBER OF SEQUENCES: 45
     CORRESPONDENCE ADDRESS:
       ADDRESSEE: Genentech, Inc.
       STREET: 460 Point San Bruno Blvd
      CITY: South San Francisco
      STATE: California
      COUNTRY: USA
      ZIP: 94080
     COMPUTER READABLE FORM:
       MEDIUM TYPE: 5.25 inch, 360 Kb floppy disk
       COMPUTER: IBM PC compatible
      OPERATING SYSTEM: PC-DOS/MS-DOS
       SOFTWARE: patin (Genentech)
     CURRENT APPLICATION DATA:
      APPLICATION NUMBER: PCT/US95/04228
      FILING DATE:
      CLASSIFICATION:
    PRIOR APPLICATION DATA:
      APPLICATION NUMBER: 08/222616
       FILING DATE: 04-APR-1994
    ATTORNEY/AGENT INFORMATION:
      NAME: Wendy M. Lee
      REGISTRATION NUMBER: 00,000
      REFERENCE/DOCKET NUMBER: 821P3PCT
    TELECOMMUNICATION INFORMATION:
      TELEPHONE: 415/225-1994
       TELEFAX: 415/952-9881
      TELEX: 910/371-7168
  INFORMATION FOR SEQ ID NO: 33:
    SEQUENCE CHARACTERISTICS:
       LENGTH: 1298 amino acids
       TYPE: amino acid
       TOPOLOGY: linear
PCT-US95-04228-33
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RESULT 12
US-08-874 678-32
; Sequence 32, Application US/08874678
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; Patent No. 5952199
  GENERAL INFORMATION:
    APPLICANT
                Davis-Smyth, Terri L.
    APPLICANT: Chen, Helen H.
APPLICANT: Aresta, Leonard
APPLICANT: Ferrara, Napoleone
    TITLE OF INVENTION: NOVEL INHIBITORS OF VASCULAR ENDOTHELIAL GROWTH FACTOR TITLE OF INVENTION: ACTIVITY, THEIR USES AND PROCESSES FOR THEIR PRODUCTION
    NUMBER OF SEQUENCES
                           48
    CORRESPONDENCE ADDRESS:
      ADDRESSEE: Flehr, Hobbach, Test, Albritton & Herbert
      STREET: Four Embarcadero Center, Suite 3400
     CITY: San Francisco
      STATE: California
      COUNTRY: United States
      ZIP: 94111-4187
   COMPUTER READABLE FORM:
      MEDIUM TYPE: Floppy disk
      COMPUTER: IBM PC compatible
      OPERATING SYSTEM: PC-DOS/MS-DOS
       SOFTWARE: PatentIn Release #1.0, Version #1.30
    CURRENT APPLICATION DATA:
      APPLICATION NUMBER: US/08/874,678
      FILING DATE: HEREWITH
       CLASSIFICATION: 435
    PRIOR APPLICATION DATA:
      APPLICATION NUMBER: US 08/643,839
       FILING DATE: 07-MAY-1996
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ATTORNEY/AGENT INFORMATION:
      NAME: Dreger, Walter H.
      REGISTRATION NUMBER: 24,190
      REFERENCE/DOCKET NUMBER: A-63291-1/WHD
    TELECOMMUNICATION INFORMATION:
      TELEPHONE: (415) 781-1989
      TELEFAX: (415) 398-3249
      TELEX: 910 277299
  INFORMATION FOR SEQ ID NO: 32:
    SEQUENCE CHARACTERISTICS:
      LENGTH: 1363 amino acids
      TYPE: amino acid
      STRANDEDNESS: unknown
      TOPOLOGY: unknown
    MOLECULE TYPE: protein
US-08-874-678-32
 Query Match 99.1%; Score 6798; DB 2; Length 1363; Best Local Similarity 99.3%; Pred. No. 0;
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 Matches 1288; Conservative
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                                                             Sequence 964, App
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                                                             Sequence 970, App
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14 US-10-081-126-2

US-10-105-901-32

US-09-375-248-19

US-10-661-740-19

US-10-105-901-33

US-10-440-464-129

US-10-105-901-3

16 US-10-473-127-949

14 US-10-105-901-34

16 US-10-473-127-951

16 US-10-783-528-61

1356 14 US-10-090-183-2

1356 15 US-10-394-322A-66

1356 17 US-10-872-198-115

1356 17 US-10-741-600-1469

Sequence 2, Appli

Sequence 32, Appl

Sequence 949, App

Sequence 34, Appl Sequence 19, Appl

Sequence 19, Appl

Sequence 951, App

Sequence 33, Appl

Sequence 3, Appli

Sequence 2, Appli

Sequence 66, Appl

Sequence 129, App

Sequence 61, Appl

Sequence 115, App

Sequence 1469, Ap

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6808

6798

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88.6

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RESULT 1
US-09-982-610-33
; Sequence 33, Application US/09982610
; Patent No. US20020146420A1
    GENERAL INFORMATION:
        APPLICANT: Genentech, Inc.
                    Bennett, Brian D.
                    Goeddel, David
                    Lee, James M.
                    Matthews, William
                    Tsai, Siao Ping
                    Wood, William I.
         TITLE OF INVENTION: PROTEIN TYROSINE KINASE AGONIST ANTIBODIES
         NUMBER OF SEQUENCES: 45
         CORRESPONDENCE ADDRESS:
              ADDRESSEE: Genentech, Inc.
              STREET: 460 Point San Bruno Blvd
              CITY: South San Francisco
              STATE: California
              COUNTRY: USA
              ZIP: 94080
         COMPUTER READABLE FORM:
              MEDIUM TYPE: 3.5 inch, 1.44 Mb floppy disk
              COMPUTER: IBM PC compatible
              OPERATING SYSTEM: PC-DOS/MS-DOS
              SOFTWARE: WinPatin (Genentech)
         CURRENT APPLICATION DATA:
             APPLICATION NUMBER: US/09/982,610
              FILING DATE: 17-Oct-2001
              CLASSIFICATION: <Unknown>
         PRIOR APPLICATION DATA:
              APPLICATION NUMBER: 08/446,648
              FILING DATE: 1996-MAY-23
              APPLICATION NUMBER: 08/222616
              FILING DATE: 04-APR-1994
         ATTORNEY/AGENT INFORMATION:
              NAME: Lee, Wendy M.
              REGISTRATION NUMBER: 40,378
              REFERENCE/DOCKET NUMBER: P0821P3PCT
         TELECOMMUNICATION INFORMATION:
              TELEPHONE: 415/225-1994
              TELEFAX: 415/952-9881
              TELEX: 910/371-7168
    INFORMATION FOR SEQ ID NO: 33:
         SEQUENCE CHARACTERISTICS:
              LENGTH: 1298 amino acids
              TYPE: Amino Acid
              TOPOLOGY: Linear
         SEQUENCE DESCRIPTION: SEQ ID NO: 33:
US-09-982-610-33
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; Sequence 67, Application US/10394322A
; Publication No. US20030232391A1
; GENERAL INFORMATION:
; APPLICANT: SUNESIS PHARMACEUTICALS, INC.
; APPLICANT: Prescott, John C.
; TITLE OF INVENTION: IDENTIFICATION OF KINASE INHIBITORS
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1356 17 US-10-741-600-1471

1356 17 US-10-926-806-10

Sequence 1471, Ap

Sequence 10, Appl

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; FILE REFERENCE: 39750-0006 US
; CURRENT APPLICATION NUMBER: US/10/394,322A
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  PRIOR APPLICATION NUMBER: US 60/366,892
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; Sequence 960, Application US/10473127 ; Publication No. US20040236091A1
; GENERAL INFORMATION:
; APPLICANT: Zycos Inc.
   TITLE OF INVENTION: TRANSLATIONAL PROFILING
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; CURRENT APPLICATION NUMBER: US/10/473,127
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PRIOR APPLICATION NUMBER: 60/279,495
PRIOR FILING DATE: 2001-03-28
                                                        2-28-01
  PRIOR APPLICATION NUMBER: 60/292,544
PRIOR FILING DATE: 2001-05-21
  PRIOR APPLICATION NUMBER: 60/310,801
; PRIOR FILING DATE: 2001-08-08
; PRIOR APPLICATION NUMBER: 60/326,370
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; PRIOR APPLICATION NUMBER: 60/358,985
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31	6808	94.4	1298	6	ABU04304	Abu04304 Human exp
32	6808	94.4	1298	6	ABU04277	Abu04277 Human exp
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     15-JAN-2001 (first entry)
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     Human Flt4 receptor tyrosine kinase long form protein sequence.
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     Human; Flt4 receptor tyrosine kinase; antibody; extracellular domain;
     lymphatic vessel detection; lymphatic tissue; lymph node tissue; endothelial venule; diagnosis; lymphoma; long form.
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KW
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     14-NOV-1994;
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XX
     (ORIN ) ORION CORP.
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PA
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     Alitalo K, Aprelikova O, Armstrong E, Korhonen J, Kaipainen A;
ΡI
     Matikainien M, Pajusola K;
XX
     WPI; 2000-571323/53.
DR
     N-PSDB; AAA37816.
DR
XX
PT
     Antibody to extracellular domain of or to an epitope unique to a
PT
     vertebrate Flt4 receptor tyrosine kinase protein useful for diagnosing
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     lymphoma and imaging lymphatic vessels or high endothelial venules in
PT
     tissue.
XX
PS
     Claim 21; Col 55-66; 66pp; English.
XX
CC
     This sequence represents the human Flt4 receptor tyrosine kinase long
     form protein. The invention relates to an antibody (I) specific to the
CC
CC
     extracellular domain of or to an epitope unique to a vertebrate Flt4
CC
     receptor tyrosine kinase protein (II). A composition comprising (I) is
     useful for detecting lymphatic vessels, lymphatic tissue comprising lymph
CC
CC
     node tissue or high endothelial venules in an organism preferably mammal
CC
     especially human. The method comprises administering the composition and
CC
     detecting (I) bound to lymphatic vessels, lymphatic tissue or high
CC
     endothelial venules. (I) is also useful for screening a biological sample
CC
     for the presence of (II) or diagnosing a disease state. The diagnosing
CC
     method of the disease state preferably lymphoma comprises obtaining a
CC
     tissue sample on a vertebrate organism suspected of being in a diseased
CC
     state characterised by in Flt4 expression in lymphatic cells or high
     endothelial venules and screening the diseased state utilising (I). (I)
CC
CC
     is also useful for imaging lymphatic vessels or high endothelial venules
CC
     in a tissue by contacting the tissue with (I) and imaging the vessels by
CC
     detecting (I) bound to the tissues
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     Protein tyrosine-kinase; SAL-S1; cell growth; differentiation.
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     04-APR-1995;
                    95WO-US004228.
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     04-APR-1994;
                    94US-00222616.
PR
xx
     (GETH ) GENENTECH INC.
PA
XX
     Bennett BD, Goeddel D, Lee JM, Matthews W, Tsai SP, Wood WI;
PΙ
XX
DR
     WPI; 1995-366160/47.
     N-PSDB; AAT03090.
DR
XX
     Agonist antibodies which activate specific protein tyrosine kinase(s) -
PT
PT
     also activate chimeric proteins of kinase extracellular domain and Ig
     constant domain, useful for studying, and therapeutic modulation of, cell
PT
PT
     growth and differentiation.
XX
PS
     Disclosure; Fig 15A-F; 125pp; English.
XX
     DNA probes based on protein tyrosine-kinase (pTK) sequences were used to
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     screen cDNA libraries to identify novel pTK genes. The SAL-S1 gene was
CC
     isolated from several megakaryocytic cell line libraries. The gene (see
CC
     AAT03090) encodes a protein (AAR85937) showing homology to FLT/FLK family
CC
CC
     pTKs. SAL-S1 can be used to target proteins in drug design, as an
CC
     (ant) agonist of other pTKs, to screen pTK ligands or to modulate
CC
     megakaryocyte/platelet adhesion
XX
SO
     Sequence 1298 AA;
                          95.0%; Score 6852; DB 2; Length 1298; 100.0%; Pred. No. 0;
  Query Match
  Best Local Similarity
  Matches 1297; Conservative
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Issued Patents:

SUMMARIES

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3	7211	100.0	1363	4	US-09-375-248-2	Sequence 2, Appli
4	7211	100.0	1363	4	US-09-169-079-4	Sequence 4, Appli
5	7157	99.3	1363	2	US-08-874-678-32	Sequence 32, Appl
6	7157	99.3	1363	3	US-08-643-839-32	Sequence 32, Appl
7	7157	99.3	1363	3	US-09-348-886-32	Sequence 32, Appl
8	6852	95.0	1298	1	US-08-222-616-33	Sequence 33, Appl
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13	6852	95.0	1298	4	US-09-169-079-2	Sequence 2, Appli
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17	6797.5	94.3	1368	3	US-09-348-886-34	Sequence 34, Appl
18	6320	87.6	1363	4	US-09-375-248-19	Sequence 19, Appl
19	6005.5	83.3	1362	2	US-08-874-678-33	Sequence 33, Appl
20	6005.5	83.3	1362	3	US-08-643-839-33	Sequence 33, Appl

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                                                                           Sequence 33, Appl

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; Sequence 2, Application US/09375248
; Patent No. 6764820
; GENERAL INFORMATION:
; APPLICANT: Ferrell, Robert E.; APPLICANT: Alitalo, Kari; APPLICANT: Finegold, David N.; APPLICANT: Karkkainen, Marika
   TITLE OF INVENTION: SCREENING AND THERAPY FOR LYMPHATIC DISORDERS INVOLVING
; TITLE OF INVENTION: THE FLT4 RECEPTOR TYROSINE KINASE (VEGFR-3)
; FILE REFERENCE: 28967/35255A
  CURRENT APPLICATION NUMBER: US/09/375,248
   CURRENT FILING DATE: 1999-08-16
; EARLIER APPLICATION NUMBER: PCT/US99/06133
; EARLIER FILING DATE: 1999-03-26
   NUMBER OF SEQ ID NOS: 28
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; LENGTH: 1363
    TYPE: PRT
    ORGANISM: Homo sapiens
US-09-375-248-2
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  Best Local Similarity 100.0%; Pred. No. 0;
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; Sequence 32, Application US/08874678
; Patent No. 5952199
; GENERAL INFORMATION:
; APPLICANT: Davis-Smyth, Terri L.
; APPLICANT: Chen, Helen H.
     APPLICANT: Predta, Leonard

APPLICANT: Ferrara, Napoleone

TITLE OF INVENTION: NOVEL INHIBITORS OF VASCULAR ENDOTHELIAL GROWTH FACTOR

TITLE OF INVENTION: ACTIVITY, THEIR USES AND PROCESSES FOR THEIR PRODUCTION
      NUMBER OF SEQUENCES: 48
                                                                                 T-7-96
     CORRESPONDENCE ADDRESS:
      ADDRESSEE: Flehr, Wohbach, Test, Albritton & Herbert
       STREET: Four Embarcadero Center, Suite 3400
       CITY: San Francisco
STATE: California
        COUNTRY: United States
        ZIP: 94111-4187
     COMPUTER READABLE FORM:
      MEDIUM TYPE: Floppy disk
        COMPUTER: IBM PC compatible
OPERATING SYSTEM: PC-DOS/MS-DOS
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       APPLICATION NUMBER: US/08/874,678
       FILING DATE: HEREWITH
       CLASSIFICATION: 435
     PRIOR APPLICATION DATA:
       APPLICATION NUMBER: US 08/643,839
       FILING DATE: 07-MAY-1996
     ATTORNEY/AGENT INFORMATION:
      NAME: Dreger, Walter H.
       REGISTRATION NUMBER: 24,190
       REFERENCE/DOCKET NUMBER: A-63291-1/WHD
     TELECOMMUNICATION INFORMATION:
       TELEPHONE: (415) 781-1989
       TELEFAX: (415) 398-3249
       TELEX: 910 277299
  INFORMATION FOR SEQ ID NO: 32:
     SEQUENCE CHARACTERISTICS:
       LENGTH: 1363 amino acids
       TYPE: amino acid
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       TOPOLOGY: unknown
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US-08-874-678-32
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US-08-222-616-33
; Sequence 33, Application US/08222616
; Patent No. 5635177
  GENERAL INFORMATION:
    APPLICANT: Bennett, Brian D.
APPLICANT: Goeddel, David
APPLICANT: Lee, James M.
    APPLICANT: Matthews, William APPLICANT: Tsai, Siao Ping APPLICANT: Wood, William I.
    TITLE OF INVENTION: PROTEIN TYROSINE KINASE AGONIST TITLE OF INVENTION: ANTIBODIES
NUMBER OF SEQUENCES: 42
     CORRESPONDENCE ADDRESS:
      ADDRESSEE: Genentech, Inc.
       STREET: 460 Point San Bruno Blvd
      CITY: South San Francisco
      STATE: California
       COUNTRY: USA
      ZIP: 94080
     COMPUTER READABLE FORM:
       MEDIUM TYPE: 5.25 inch, 360 Kb floppy disk
       COMPUTER: IBM PC compatible
       OPERATING SYSTEM: PC-DOS/MS-DOS
       SOFTWARE: patin (Genentech)
     CURRENT APPLICATION DATA:
       APPLICATION NUMBER: US/08/222,616
       FILING DATE: 4-APR-1994
       CLASSIFICATION: 530
     PRIOR APPLICATION DATA:
      APPLICATION NUMBER: PCT/US93/00586
       FILING DATE: 22-JAN-1993
     PRIOR APPLICATION DATA:
       APPLICATION NUMBER: 07/826935
       FILING DATE: 22-JAN-1992
    ATTORNEY/AGENT INFORMATION:
      NAME: Lee, Wendy M.
      REGISTRATION NUMBER:
      REFERENCE/DOCKET NUMBER: 821P2
    TELECOMMUNICATION INFORMATION:
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TELEPHONE: 415/225-1994
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       TYPE: amino acid
       TOPOLOGY: linear
US-08-222-616-33
                         95.0%; Score 6852; DB 1; Length 1298;
  Ouery Match
  Best Local Similarity 100.0%; Pred. No. 0;
  Matches 1297; Conservative 0; Mismatches
                                                  0; Indels
RESULT 11
US-08-446-648-33
; Sequence 33, Application US/08446648
; Patent No. 6331302
  GENERAL INFORMATION:
    APPLICANT: Genentech, Inc.
     APPLICANT: Bennett, Brian D.
    APPLICANT: Goeddel, David
    APPLICANT: Lee, James M.
    APPLICANT: Matthews, William
     APPLICANT: Tsai, Siao Ping
    APPLICANT: Wood, William I.
     TITLE OF INVENTION: PROTEIN TYROSINE KINASE AGONIST ANTIBODIES
    NUMBER OF SEQUENCES: 45
    CORRESPONDENCE ADDRESS:
      ADDRESSEE: Genentech, Inc.
      STREET: 460 Point San Bruno Blvd
      CITY: South San Francisco
STATE: California
      COUNTRY: USA
      ZIP: 94080
     COMPUTER READABLE FORM:
      MEDIUM TYPE: 3.5 inch, 1.44 Mb floppy disk
       COMPUTER: IBM PC compatible
      OPERATING SYSTEM: PC-DOS/MS-DOS
      SOFTWARE: WinPatin (Genentech)
    CURRENT APPLICATION DATA:
      APPLICATION NUMBER: US/08/446,648
      FILING DATE:
      CLASSIFICATION: 435
    PRIOR APPLICATION DATA:
      APPLICATION NUMBER: 08/222616
      FILING DATE: 04-APR-1994
    ATTORNEY/AGENT INFORMATION:
      NAME: Lee, Wendy M.
      REGISTRATION NUMBER:
                            40,378
      REFERENCE/DOCKET NUMBER: P0821P3PCT
    TELECOMMUNICATION INFORMATION:
      TELEPHONE: 415/225-1994
      TELEFAX: 415/952-9881
      TELEX: 910/371-7168
  INFORMATION FOR SEQ ID NO: 33:
    SEQUENCE CHARACTERISTICS:
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      TYPE: Amino Acid
      TOPOLOGY: Linear
US-08-446-648-33
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PGPUBS:

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RESULT 2
US-09-765-534B-6
; Sequence 4, Application US/09765534B
; Publication No. US20040037820A1
; GENERAL INFORMATION:
; APPLICANT: Alitalo, Kari
; APPLICANT: Kaipainen, Arja
; APPLICANT: Valtola, Reija
; APPLICANT: Jussila, Lotta
; TITLE OF INVENTION: Flt4 (VEGFR-3) as a Target for Tumor Imaging and Anti-Tumor Therapy
; FILE REFERENCE: 28113/34891
; CURRENT APPLICATION NUMBER: US/09/765,534B
; CURRENT FILING DATE: 2001-01-19
; PRIOR APPLICATION NUMBER: 08/901,710
; PRIOR FILING DATE: 1997-07-28
; PRIOR APPLICATION NUMBER: 08/340,011
; PRIOR FILING DATE: 1994-11-14
; PRIOR APPLICATION NUMBER: 08/257,754
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3	4919.5	68.2	1379	2	JC4954	vascular endotheli
4	2804	38.9	1348	2	S51656	vascular endotheli
5	2753	38.2	1356	2	JC1402	protein-tyrosine k
6	2669	37.0	1367	2	A41228	protein-tyrosine k
7	2469	34.2	1333	2	178875	receptor tyrosine
8	2463	34.2	1338	2	S09982	protein-tyrosine k
9	2444.5	33.9	1336	2	160598	Fit-1 tyrosine kin
10	2437.5	33.8	1330	2	S49010	embryonic receptor
11	1283.5	17.8	367	2	B56598	endothelial kinase
12	1163.5	16.1	1089	1	PFHUGA	platelet-derived g
13	1151	16.0	1088	1	PFRTGA	platelet-derived g
14	1125	15.6	1089	1	S33727	platelet-derived g
15	1122.5	15.6	1087	2	151552	platelet-derived g
16	1084	15.0	1106	1	PFHUGB	platelet-derived g
17	1052.5	14.6	1098	1	PFMSRB	platelet-derived g
18	1051	14.6	975	2	T30816	macrophage colony-
19	1044	14.5	978	1	A49814	protein-tyrosine k
20	1042.5	14.5	976	1	TVHUKT	protein-tyrosine k
21	1033.5	14.3	975	1	TVMSKT	protein-tyrosine k
22	1024	14.2	977	2	I45877	protein-tyrosine k
23	1021.5	14.2	960	1	JN0677	protein-tyrosine k
24	1018	14.1	954	2	I51703	c-kit-related kina
25	1012	14.0	1048	2	T30815	platelet-derived g
26	967	13.4	980	1	TVCTMD	macrophage colony-
27	956	13.3	941	1	TVMVMD	protein-tyrosine k
28	954.5	13.2	972	1	TVHUMD	macrophage colony-
29	954.5	13.2	992	2	A39931	protein-tyrosine k
30	951	13.2	1000	2	S18827	Flt3 protein - mou
31	949	13.2	978	2	S16385	macrophage colony-
32	930.5	12.9	976	1	TVMSMD	macrophage colony-
33	896	12.4	806	2	A35963	protein-tyrosine k
34	895.5	12.4	687	2	A49636	soluble vascular e
35	893.5	12.4	993	2	A36873	protein-tyrosine k
36	886.5	12.3	797	2	S38579	fibroblast growth
37	873	12.1	790	1	FOMVHZ	gag-kit polyprotei
38	869	12.1	822	1	TVHUFG	fibroblast growth
39	866	12.0	822	1	TVMSFG	fibroblast growth
40	865	12.0	814	1	A39752	fibroblast growth
41	863.5	12.0	822	2	B54846	fibroblast growth
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44	861.5	11.9	800	1	TVHU2F	fibroblast growth
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N; Alternate names: class III receptor tyrosine kinase FLT4; fms-like tyrosine kinase 4
C; Species: Homo sapiens (man)
C;Date: 21-Jan-1994 #sequence_revision 18-Nov-1994 #text_change 09-Jul-2004
C; Accession: A48999; A44930; G02316; S36130; A42010
R; Pajusola, K.; Aprelikova, O.; Korhonen, J.; Kaipainen, A.; Pertovaara, L.; Alitalo, R.;
Alitalo, K.
Cancer Res. 52, 5738-5743, 1992
A; Title: FLT4 receptor tyrosine kinase contains seven immunoglobulin-like loops and is
expressed in multiple human tissues and cell lines.
A; Reference number: A48999; MUID: 93007958; PMID: 1327515
A:Accession: A48999
A; Status: preliminary; not compared with conceptual translation
A; Molecule type: mRNA
A; Residues: 1-1298 < PAJ>
A;Cross-references: UNIPROT:P35916; PIDN:AAB23636.1; PID:g257352
A; Experimental source: HEL erythroleukemia cells
A; Note: sequence extracted from NCBI backbone (NCBIP:115335)
R; Aprelikova, O.; Pajusola, K.; Partanen, J.; Armstrong, E.; Alitalo, R.; Bailey, S.K.;
McMahon, J.; Wasmuth, J.; Huebner, K.; Alitalo, K.
Cancer Res. 52, 746-748, 1992
A; Title: FLT4, a novel class III receptor tyrosine kinase in chromosome 5q33-qter.
A; Reference number: A44930; MUID: 92119639; PMID: 1310071
A; Accession: A44930
A; Status: preliminary; not compared with conceptual translation
A; Molecule type: mRNA
A; Residues: 761-1190 < APR>
A; Cross-references: GB:X68203; NID:g31433
A; Note: sequence extracted from NCBI backbone (NCBIP:78155)
R; Wood, W.I.
submitted to the EMBL Data Library, December 1995
A; Reference number: H01039
A; Accession: G02316
A; Status: translated from GB/EMBL/DDBJ
A; Molecule type: mRNA
A; Residues: 1-1298 < WOO>
A; Cross-references: EMBL: U43143; NID: g1150990; PIDN: AAA85215.1; PID: g1150991
submitted to the EMBL Data Library, December 1992
A; Reference number: S36130
A; Accession: S36130
A; Molecule type: mRNA
A; Residues: 1-23, 'D', 25-744, 'P', 746-751, 'RP', 754-889, 'Q', 891-1127, 'V', 1129-1145, 'H', 1147-
1163, 'D', 1165-1298 <GAL1>
A;Cross-references: EMBL:X69878; NID:g297049; PIDN:CAA49505.1; PID:g297050
R;Galland, F.; Karamysheva, A.; Mattei, M.G.; Rosnet, O.; Marchetto, S.; Birnbaum, D.
Genomics 13, 475-478, 1992
A; Title: Chromosomal localization of FLT4, a novel receptor-type tyrosine kinase gene.
A; Reference number: A42010; MUID: 92307693; PMID: 1319394
A; Accession: A42010
A; Status: preliminary; nucleic acid sequence not shown; not compared with conceptual
translation
A; Molecule type: mRNA
A; Residues: 776-889, 'Q', 891-1127, 'V', 1129-1145, 'H', 1147-1163, 'D', 1165-1200 <GAL2>
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A;Gene: GDB:FLT4
A; Cross-references: GDB:128732; OMIM:136352
A; Map position: 5q34-5q35
C; Superfamily: unassigned Ser/Thr or Tyr-specific protein kinases; protein kinase
homology
C; Keywords: ATP; autophosphorylation; phosphoprotein; phosphotransferase; transmembrane
protein; tyrosine-specific protein kinase
F;1-23/Domain: signal sequence #status predicted <SIG>
F;24-1298/Product: protein-tyrosine kinase FLT4 #status predicted <MAT>
F;843-1176/Domain: protein kinase homology <KIN>
F;851-859/Region: protein kinase ATP-binding motif
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  Best Local Similarity 100.0%; Pred. No. 2.3e-301;
Matches 1297; Conservative
                                 0; Mismatches
                                                  0; Indels
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Qy Db		EGTTAASSYVFVRDFEQPFINKPDTLLVNRKDAMWVPCLVSIPGLNVTLRSQSSVLWPDG	
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C; Species: Mus musculus (house mouse)
C;Date: 26-Jul-1996 #sequence revision 26-Jul-1996 #text change 16-Aug-2004
C; Accession: I58375; B42010
R; Finnerty, H.; Kelleher, K.; Morris, G.E.; Bean, K.; Merberg, D.M.; Kriz, R.; Morris,
J.C.; Sookdeo, H.; Turner, K.J.; Wood, C.R.
Oncogene 8, 2293-2298, 1993
A; Title: Molecular cloning of murine FLT and FLT4.
A; Reference number: I58375; MUID: 93330572; PMID: 8393164
A:Accession: I58375
A; Status: preliminary; translated from GB/EMBL/DDBJ
A; Molecule type: mRNA
A; Residues: 1-1363 < RES>
A; Cross-references: UNIPROT: P35917; GB:L07296; NID: g293780; PIDN: AAA40077.1; PID: g293781
R;Galland, F.; Karamysheva, A.; Mattei, M.G.; Rosnet, O.; Marchetto, S.; Birnbaum, D.
Genomics 13, 475-478, 1992
A; Title: Chromosomal localization of FLT4, a novel receptor-type tyrosine kinase gene.
A; Reference number: A42010; MUID: 92307693; PMID: 1319394
A; Accession: B42010
A; Status: preliminary; nucleic acid sequence not shown; not compared with conceptual
translation
A; Molecule type: DNA
A; Residues: 1033-1072 < GAL>
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A:Gene: FTL4
C; Superfamily: protein kinase homology
C; Keywords: ATP; phosphotransferase; tyrosine-specific protein kinase
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RESILT 5
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protein-tyrosine kinase (EC 2.7.1.112) KDR - human
C; Species: Homo sapiens (man)
C;Date: 30-Sep-1993 #sequence revision 28-Aug-1998 #text_change 16-Aug-2004
C; Accession: JC1402; I58357
R; Terman, B.I.; Dougher-Vermazen, M.; Carrion, M.E.; Dimitrov, D.; Armellino, D.C.;
Gospodarowicz, D.; Boehlen, P.
Biochem. Biophys. Res. Commun. 187, 1579-1586, 1992
A; Title: Identification of the KDR tyrosine kinase as a receptor for vascular endothelial
cell growth factor.
A; Reference number: JC1402; MUID: 93038639; PMID: 1417831
A; Accession: JC1402
A; Status: nucleic acid sequence not shown
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A; Residues: 1-48,'I',50-616,'E',618-700,'E',702-748,'F',750-1263,'I',1265-1290,'P',1292-
A; Cross-references: UNIPROT: P35968; EMBL: X61656; NID:q31717
R; Terman, B.I.; Carrion, M.E.; Kovacs, E.; Rasmussen, B.A.; Eddy, R.L.; Shows, T.B.
Oncogene 6, 1677-1683, 1991
A; Title: Identification of a new endothelial cell growth factor receptor tyrosine kinase.
A; Reference number: I58357; MUID: 92019839; PMID: 1656371
A: Accession: I58357
A; Status: preliminary; translated from GB/EMBL/DDBJ
A; Molecule type: mRNA
A; Residues: 3-1356 <TER2>
A; Cross-references: GB:L04947; NID:q186674; PIDN:AAA59459.1; PID:q186675
C; Comment: This enzyme acts as a receptor for vascular endothelial cell growth factor.
C:Genetics:
A; Gene: GDB: KDR
A; Cross-references: GDB:127921; OMIM:191306
A; Map position: 4q12-4q12
C; Superfamily: protein kinase homology
C; Keywords: ATP; phosphotransferase; transmembrane protein; tyrosine-specific protein
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F;766-790/Domain: transmembrane #status predicted <TMM>
F;832-1167/Domain: protein kinase homology <KIN>
F;840-848/Region: protein kinase ATP-binding motif
 Query Match 38.2%; Score 2753; DB 2; Length 1356; Best Local Similarity 43.9%; Pred. No. 1.1e-116;
  Matches 606; Conservative 205; Mismatches 480; Indels 88; Gaps
Qy
            1 MQRGAALCLRLWLCLGLLDGLVS--GYSMTPPTLNITEESHVIDTGDSLSISCRGQHPLE 58
             1: | : | | | | :
                                      1
           {\tt 1} {\tt MESKVLLAVALWLCVETRAASVGLPSVSLDLPRLSIQKDILTIKANTTLQITCRGQRDLD} {\tt 60} \\
Db
          59 WAWPGAQEAPATGDKDSEDTGVVRDC-EGTDARPYCKVLLLHEVHANDTGSYVCYYKYIK 117
Qу
                       11
                                   | :| :|
                                              - :|| | : :| ||||:| |:|:
          61 WLWPNNQSG-----SEQRVEVTECSDGL----FCKTLTIPKVIGNDTGAYKCFYR--- 106
Db
          118 ARIEGTTAASSYVFVRDFEQPFINKPDT-----LLVNRKDAMWVPCLVSIPGLNVTL-- 169
Oy
                | |: ||:|:|: |||
                                           . : |: ::||| || || || ||
             ---ETDLASVIYVYVQDYRSPFIASVSDQHGVVYITENKNKTVVIPCLGSISNLNVSLCA 163
Db
Ov
          170 RSQSSVLWPDGQEVVWDDRRGMLVSTPLLHDALYLQCETTWGDQDFLSNPFLVHITGNEL 229
                     Db
          164 RYPEKRFVPDGNRISWDSKKGFTIPSYMISYAGMVFCEAKINDESYQSIMYIVVVVGYRI 223
Qу
         230 YDIQLLPRKSLELLVGEKLVLNCTVWAEFNSGVTFDWDYPGKQAERGKWVPERRSQQTHT 289
                      ||\cdot||\cdot||
          224 YDVVLSPSHGIELSVGEKLVLNCTARTELNVGIDFNWEYPSSKHQHKKLVNRDLKTQSGS 283
Db
Οv
         290 ELS---SILTIHNVSOHDLGSYVCKANNGIORFRESTEVIVHENPFISVEWLKGPILEAT 346
```

	Db	284	: :: :: : :	43
	Qy	347	AGDELVKLPVKLAAYPPPEFQWYKDGKALSGRHSPHALVLKEVTEASTGTYTLALWN 40	03
	Db	. 344	VG-ERVRIPAKYLGYPPPEIKWYKNGIPLESNHTIKAGHVLTIMEVSERDTGNYTVILTN 40	02
	Qy	404	SAAGLRRNISLELVVNVPPQIHEKEASSP-SIYSRHSRQALTCTAYGVPLPLSIQWHWRP 46 : ::: :	62
	Db	403	PISKEKQSHVVSLVVYVPPQIGEKSLISPVDSYQYGTTQTLTCTVYAIPPPHHIHWYWQL 40	62
	Qy	463	WTPCKMFAQRSLRRRQQQDLMPQCRDWRAVTTQDAVNPIESLDTWTEFVEGKNKTVSKLV 5:	22
	Db	463	EEECANEPSQAVSVTNPYP-CEEWRSVEDFQGGNKIEVNKNQFALIEGKNKTVSTLV 5:	18
	Qу	523	IQNANVSAMYKCVVSNKVGQDERLIYFYVTTIPDGFTIESKPSEELLEGQPVLLSCQADS 58	82
	Db	519	IQAANVSALYKCEAVNKVGRGERVISFHVTRGPEITLQPDMQPTEQESVSLWCTADR 5	75
	Qу	583	YKYEHLRWYRLNLSTLHDAHGNPLLLDCKNVHLFATPLAASLEEVAPGARHATLSL 6:	38
	Db		STFENLTWYKLGPQPLPIHVGELPTPVCKNLDTLWKLNATMFSNSTNDILIM 62	
	Qy		SIPRVAPEHEGHYVCEVQDRRSHDKHCHKKYLSVQALEAPRLTQNLTDLLVNVSDSLEMQ 6: : : : : : : :	
-	Db		ELKNASLQDQGDYVCLAQDRKTKKRHCVVRQLTVLERVAPTITGNLENQTTSIGESIEVS 68	
	Qy		CLVAGAHAPSIVWYKDERLLEEKSGVDLADSNQKLSIQRVREEDAGRYLCSVCNAKGCVN 79	
	Db		CTASGNPPPQIMWFKDNETLVEDSGIVLKDGNRNLTIRRVRKEDEGLYTCQACSVLGCAK 74	
	Qy Db		SSASVAVEGSEDKGSMEIVILVGTGVIAVFFWVLLLLIFCNMRRPAHADIKTGYLSIIMD 8: : ::: :: : : : :: ::	
	Db		VEAFFIIEGAQEKTNLEIIILVGTTVIAMFFWLLLVIILGTVKRANGGELKTGYLSIVMD 80 PGEVPLEEQCEYLSYDASQWEFPRERLHLGRVLGYGAFGKVVEASAFGIHKGSSCDTVAV 8	
	Qy Db		: : : : : : :: PDELPLDEHCERLPYDASKWEFPRDRLNLGKPLGRGAFGQEIEADAFGIDKTATCRTVAV 86	
	Qy		KMLKEGATASEHRALMSELKILIHIGNHLNVVNLLGACTKPQGPLMVIVEFCKYGNLSNF 93	
	Db			
	Qу	939	LRAKRDAFSPCAEKSPEQRGRFRAMVELARLDRRRPGSSDRVLFARFSKTEGGARRA 99	95
	Db	928	: :	81 ·
	Qу	996	SPDQEA-EDLWLSPLTMEDLVCYSFQVARGMEFLASRKCIHRDLAARNILLSESDV 10	050
	Db	982	:: : : : : : : : :	041
	Qy	1051	VKICDFGLARDIYKDPDYVRKGSARLPLKWMAPESIFDKVYTTQSDVWSFGVLLWEIFSL 1	110
	Db	1042		101
	Qу	1111	GASPYPGVQINEEFCQRLRDGTRMRAPELATPAIRRIMLNCWSGDPKARPAFSELVEILG 1	170
	Db	1102	: : : : : : : :	161
	Qу	1171	DLLQGRGLQEEEEVCMAPRS-SQSSEEGSFSQVSTMALHIAQADAEDSPPSLQRHSLAAR 12:	229
	Db	1162	NLLQANAQQDGKDYIVLPISETLSMEEDSGLSLPTSPVSCMEEEEVCDPKFH 12	213
	Qy	1230	YYNWVSFPGCLARGAETRGSSRMKTFEEFPM-TPTTYKGSVDNQTDSGMVLASEEFEQIE 12	288
	Db	1214	YDNTAGISQYLQNSKRKSRPVSVKTFEDIPLEEPEVKVIPDDNQTDSGMVLASEELKTLE 12	273
	Qу	1289	SRHRQESGFSCKGPGQNVAVTRAHPDSQGRRRRPERGARGGQVFYNSEYGELSEPSEED 134	17
	Db	1274	DRTKLSPSFGGMVPSKSRESVASEGSNQTSGYQSGYHSDDTDTTVYSSEE 132	23

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protein-tyrosine kinase (EC 2.7.1.112) flt1 precursor - human
N; Alternate names: receptor-type tyrosine kinase flt
C; Species: Homo sapiens (man)
C;Date: 12-Feb-1993 #sequence_revision 12-Feb-1993 #text_change 16-Aug-2004
C; Accession: S09982
R;Shibuya, M.; Yamaguchi, S.; Yamane, A.; Ikeda, T.; Tojo, A.; Matsushime, H.; Sato, M.
Oncogene 5, 519-524, 1990
A; Title: Nucleotide sequence and expression of a novel human receptor-type tyrosine
kinase gene (flt) closely related to the fms family.
A; Reference number: S09982; MUID: 90221591; PMID: 2158038
A; Accession: S09982
A; Molecule type: mRNA
A; Residues: 1-1338 <SHI>
A;Cross-references: UNIPROT:P17948; EMBL:X51602; NID:g31431; PIDN:CAA35946.1; PID:g31432
C:Genetics:
A;Gene: GDB:FLT1
A; Cross-references: GDB:120616; OMIM:165070
A; Map position: 13q12-13q12
C; Superfamily: protein kinase homology
C; Keywords: ATP; autophosphorylation; glycoprotein; membrane protein; phosphoprotein;
phosphotransferase; tyrosine-specific protein kinase
F;1-22/Domain: signal sequence #status predicted <SIG>
F;23-1338/Product: protein-tyrosine kinase flt #status predicted <MAT>
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F;833-841/Region: protein kinase ATP-binding motif
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   Query Match
   Best Local Similarity 39.9%; Pred. No. 1.2e-103;
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Qy
                     Db
                  8 GVLLC-ALLSCL-LLTGSSSGSKLKDPELSLKGTQHIMQAGQTLHLQCRGEAAHKWS--- 62
                64 AQEAPATGDKDSEDTGVVRDCEGTDARPYCKVLLLHEVHANDTGSYVCYYKYIKARIEGT 123
Qy
                                 Db
                63 ---LPEMVSKESERLSITKSACGRNGKQFCSTLTLNTAQANHTGFYSCKYLAVPTSKKKE 119
               124 TAASSYVFVRDFEQPFINK----PDTLLVNRKDAMWVPCLVSIPGLNVTLRS-QSSVLWP 178
Qу
                                                     | :: |:|: | :||:
               120 TESAIYIFISDTGRPFVEMYSEIPEIIHMTEGRELVIPCRVTSPNITVTLKKFPLDTLIP 179
Db
Qy
               179 DGQEVVWDDRRGMLVSTPLLHDALYLQCETTWGDQDFLSNPFLVHITGNELYDIQLLPRK 238
               ||:::|| |:| ::| : | || | ::| :| :| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::| ::|
Db
               239 SLELLVGEKLVLNCTVWAEFNSGVTFDWDYPGKQAERGKWVPERRSQQTHTELS---SIL 295
Qу
                      239 PVKLLRGHTLVLNCTATTPLNTRVOMTWSYPDEKNKRAS--VRRRIDOSNSHANIFYSVL 296
Db
Qу
               296 TIHNVSQHDLGSYVCKANNGIQRFRESTEVIVHENPFISVEWLKGPILEATAGDELVKLP 355
                     297 TIDKMONKDKGLYTCRVRSGPSFKSVNTSVHIYDKAFITVKHRKQQVLETVAGKRSYRLS 356
               356 VKLAAYPPPEFQWYKDGKALSGRHS-----PHALVLKEVTEASTGTYTLALWNSAAGLRR 410
Qу
                     ::|::|:[|| | | ||: |
Db
               357 MKVKAFPSPEVVWLKDGLPATEKSARYLTRGYSLIIKDVTEEDAGNYTILLSIKQSNVFK 416
Qу
               411 NISLELVVNVPPQIHEKEASS---PSIYSRHSRQALTCTAYGVPLPLSIQWHWRPWTPCK 467
                     417 NLTATLIVNVKPQIYEKAVSSFPDPALYPLGSRQILTCTAYGIPQP-TIKWFWHPCNHNH 475
Db
               468 MFAQRSLRRRQQQDLMPQCRDWRAVTTQDAVNPIESLDTWTEFVEGKNKTVSKLVIQNAN 527
Ov
                               :: : : : | [|]:
                                                                               :[][[] :::
               476 SEARCDFCSNNEESFILD------ADSNMGNRIESITQRMAIIEGKNKMASTLVVADSR 528
Db
               528 VSAMYKCVVSNKVGQDERLIYFYVTTIPDGF--TIESKPSEELLEGQPVLLSCQADSYKY 585
Qу
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Db	529	${\tt ISGIYICIASNKVGTVGRNISFYITDVPNGFHVNLEKMPTEGEDLKLSCTVNKFLY}$	584
Qy		EHLRWYRLNLSTLHDAHGNPLLLDCKNVHLFATPLAASLEEVAPGARHA-TLSLSIP : : : : : : : :	
Db	585	RDVTWILLRTVNNRTMHYSISKQKMAITKEHSITLNLTIM	624
Qу	642	RVAPEHEGHYVCEVQDRRSHDKHCHKKYLSVQALEAPRLTQNLTDLLVNVSDSLEMQCLV ::: :::: : : : :	701
Db	625	NVSLQDSGTYACRARNVYTGEEILQKKEITIRDQEAPYLLRNLSDHTVAISSSTTLDCHA	684
Qy	702	AGAHAPSIVWYKDERLLEEKSGVDLADSNQKLSIQRVREEDAGRYLCSVCNAKGCVNSSA	761
Db	685	NGVPEPQITWFKNNHKIQQEPGIILGPGSSTLFIERVTEEDEGVYHCKATNQKGSVESSA	744
Qу	762	SVAVEGSEDKGSMEIVILVGTGVIAVFFWVLLLLIFCNMRRPAHADIKTGYLSIIMDPGE : : : :	821
Db	745	YLTVQGTSDKSNLELITLTCTCVAATLFWLLLTLLIRKMKR-SSSEIKTDYLSIIMDPDE	803
Qу	822	VPLEEQCEYLSYDASQWEFPRERLHLGRVLGYGAFGKVVEASAFGIHKGSSCDTVAVKML	881
Db	804	VPLDEQCERLPYDASKWEFARERLKLGKSLGRGAFGKVVQASAFGIKKSPTCRTVAVKML	863
Qy	882	KEGATASEHRALMSELKILIHIGNHLNVVNLLGACTKPQGPLMVIVEFCKYGNLSNFLRA	941
Db	864	KEGATASEYKALMTELKILTHIGHHLNVVNLLGACTKQGGPLMVIVEYCKYGNLSNYLKS	923
Qy	942	KRDAFSPCAEKSPEQRGRFRAMVELARLDRRRPGSSDRVLFARFSKTEGGARRASPD	998
Db	924	KRDLFFLNKDAALHMEPKKEKMEPGLEQGKKPRLDSVTSSESFASSGFQEDKSLSDVEEE	983
Qy	999	QEAEDLWLSPLTMEDLVCYSFQVARGMEFLASRKCIHRDLAARNILLSESDVVKICDFGL	1058
Db .	984	EDSDGFYKEPITMEDLISYSFQVARGMEFLSSRKCIHRDLAARNILLSENNVVKICDFGL	1043
Qу	1059	ARDIYKDPDYVRKGSARLPLKWMAPESIFDKVYTTQSDVWSFGVLLWEIFSLGASPYPGV	1118
Db	1044	ARDIYKNPDYVRKGDTRLPLKWMAPESIFDKIYSTKSDVWSYGVLLWEIFSLGGSPYPGV	1103
Qy	1119	QINEEFCQRLRDGTRMRAPELATPAIRRIMLNCWSGDPKARPAFSELVEILGDLLQGRGL	1178
Db	1104	QMDEDFCSRLREGMRMRAPEYSTPEIYQIMLDCWHRDPKERPRFAELVEKLGDLLQANVQ	1163
Qy	1179	QEEEEVCMAPRSSQSSEEGSFSQVSTMALHIAQADAEDSPPSLQRHSLAARYY	1231
Db	1164	QDGKDYIPINAILTGNSGFTYSTPAFSEDFFKESISAPKFNSGSSDDVRYV	1214
Qy	1232	NWVSFPGCLARGAETRGSSRMKTFEEFPMTPTTYKGSVDNQTDSGMVLASEEFEQIE	1288
Db	1215	NAFKFMSLERIKTFEELLPNATSMFDDYQGDSSTLLASPMLKRFTWTD	1262
Qy	1289	SRHRQESGFSCKGPGQNVAVTRAHPDSQGRRR 1320	
Db	1263	SKPKASLKIDLRVTSKSKESGLSDVSRPSFCHSSCGHVSEGKRR 1306	

SUMMARIES

		8				
Result		Query				
No.	Score	Match	Length	DB	ID	Description
1	7170	99.4	1363	2	Q86W07	Q86w07 homo sapien
2	6852	95.0	1298	1	VGR3 HUMAN	P35916 homo sapien
3	6811	94.5	1298	2	Q86W08	Q86w08 homo sapien
4	6320	87.6	1363	1	VGR3 MOUSE	P35917 mus musculu
5	6278	87.1	1363	2	Q91ZT1	Q91zt1 rattus norv
6	5085	70.5	1092	2	Q91ZT0	Q91zt0 rattus norv
7	4919.5	68.2	1379	2	P79701	P79701 coturnix co
8	2804	38.9	1348	1	VGR2 COTJA	P52583 coturnix co
9	2793	38.7	1348	2	Q677M1	Q677ml gallus gall
10	2762	38.3	1356	1	VGR2 HUMAN	P35968 homo sapien

```
11
     2681
            37.2
                  1345 2 Q8VCD0
                                                      O8vcd0 mus musculu
                   1367 1 VGR2_MOUSE
                                                      P35918 mus musculu
12
     2669
            37.0
                   1343 1 VGR2 RAT
                                                      008775 rattus norv
13
     2646
            36.7
14
      2511
            34.8
                   1327
                         2
                            Q8QHL3
                                                      Q8qhl3 gallus gall
                   1333 1
                            VGR1 MOUSE
                                                     P35969 mus musculu
15
      2469
            34.2
                            VGR1_HUMAN
                                                      P17948 h vascular
16
      2463
            34.2
                   1338 1
                                                      P53767 rattus norv
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   2444.5
            33.9
                   1336
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18
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            32.0
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                            VGR2_BRARE
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1.9
     1361
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                            Q75WK4
20
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            17.8
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                            Q8AXC8
21
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                                                      P16234 homo sapien
                   1089
22
   1163.5
            16.1
                         1
     1160
                   1089
                        1
                            PGDS MOUSE
                                                      P26618 mus musculu
23
            16.1
24
     1158
            16.1
                   1089
                         2
                            O7TSJ3
                                                      Q7tsj3 mus musculu
25
     1151
            16.0
                   1088
                        1 PGDS RAT
                                                      P20786 rattus norv
                                                      Q9puf6 gallus gall
26
   1150.5
            16.0
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                            Q9PUF6
27
     1134
            15.7
                   1079 2 Q6E7G6
                                                      Q6e7g6 canis famil
                                                      Q9pul0 brachydanio
28
     1128
            15.6
                    404
                         2
                            Q9PUL0
                                                      Q8t4m0 drosophila
     1127
            15.6
                   1474
                         2
                            Q8T4M0
29
30 1123.5
                   1087
                                                      Q7zy71 xenopus lae
            15.6
                        2 Q7ZY71
   1122.5
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                         1 PGDS XENLA
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31
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                                                      Q8ipg1 drosophila
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                            Q8IPG1
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   1106.5
                                                      Q8t418 drosophila
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                        2 O8T4L8
39
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                         2
                            Q8AXC7
                                                      Q8axc7 fugu rubrip
40
            15.0
                                                      Q6qnf3 canis famil
     1084
                   1103 2
                            O6ONF3
41
     1084
                   1106 1
                            PGDR HUMAN
                                                      P09619 homo sapien
            15.0
   1071.5
                    992 2
                            0UXA8Q
                                                      Q8axu0 oncorhynchu
42
            14.9
43
   1070.5
            14.8
                    977
                         2
                            Q98SU1
                                                      Q98sul danio nigro
44
     1070
            14.8
                   1383
                         2
                            070840
                                                      Q7q840 anopheles g
45
     1065
            14.8
                    977 2 Q98SU4
                                                      Q98su4 danio albol
```

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RESULT 2
VGR3_HUMAN
     VGR3 HUMAN
                    STANDARD;
                                    PRT: 1298 AA.
ID
AC
     P35916;
DT
     01-JUN-1994 (Rel. 29, Created)
     01-OCT-1996 (Rel. 34, Last sequence update)
DТ
     25-JAN-2005 (Rel. 46, Last annotation update)
DE
     Vascular endothelial growth factor receptor 3 precursor (EC 2.7.1.112)
DΕ
     (VEGFR-3) (Tyrosine-protein kinase receptor FLT4).
GN
     Name=FLT4:
os
     Homo sapiens (Human).
OC
     Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;
OC
     Mammalia; Eutheria; Primates; Catarrhini; Hominidae; Homo.
OX
     NCBI TaxID=9606;
RN
     [1]
RP
     SEQUENCE FROM N.A., AND VARIANTS GLN-890 AND HIS-1146.
     TISSUE=Placenta;
RC
RX
     MEDLINE=93241723; PubMed=8386825;
RA
     Galland F., Karamysheva A., Pebusque M.-J., Borg J.-P., Rottapel R.,
RA
     Dubreuil P., Rosnet O., Birnbaum D.;
RT
     "The FLT4 gene encodes a transmembrane tyrosine kinase related to the
RT
     vascular endothelial growth factor receptor.";
RL
     Oncogene 8:1233-1240(1993).
RN
     [2]
RP
     SEQUENCE FROM N.A.
RX
     MEDLINE=93007958; PubMed=1327515;
RA
     Pajusola K., Aprelikova O., Korhonen J., Kaipainen A., Pertovaara L.,
RA
     Alitalo R., Alitalo K.;
RT
     "FLT4 receptor tyrosine kinase contains seven immunoglobulin-like
RT
     loops and is expressed in multiple human tissues and cell lines.";
     Cancer Res. 52:5738-5743(1992).
RL
RN
     [3]
     ERRATUM.
RΡ
```

```
Pajusola K., Aprelikova O., Korhonen J., Kaipainen A., Pertovaara L.,
RA
RÁ
     Alitalo R., Alitalo K.;
RL
     Cancer Res. 53:3845-3845(1993).
RN
RP
     SEQUENCE FROM N.A.
     Lee J., Gray A., Yuan J., Luoh S.M., Avraham H., Wood W.I.;
RA
     Submitted (JAN-1996) to the EMBL/GenBank/DDBJ databases.
RL
RN
     SEQUENCE OF 761-1190 FROM N.A.
RP
RX
     MEDLINE=92119639; PubMed=1310071;
RA
     Aprelikova O., Pajusola K., Partanen J., Armstrong E., Alitalo R.,
     Bailey S.K., McMahon J., Wasmuth J., Huebner K., Alitalo K.;
RA
RT
     "FLT4, a novel class III receptor tyrosine kinase in chromosome 5q33-
RT
     ater.";
RL
     Cancer Res. 52:746-748(1992).
RN
     [6]
RP
     SEQUENCE OF 25-39.
     PubMed=15340161; DOI=10.1110/ps.04682504;
RX
     Zhang Z., Henzel W.J.;
RA
     "Signal peptide prediction based on analysis of experimentally
RT
     verified cleavage sites.";
ŔŦ
RL
     Protein Sci. 13:2819-2824(2004).
RN
     [7]
     VARIANT HEREDITARY LYMPHEDEMA I LEU-1114.
RP
RX
     PubMed=9817924; DOI=10.1093/hmg/7.13.2073;
     Ferrell R.E., Levinson K.L., Esman J.H., Kimak M.A., Lawrence E.C.,
RA
RA
     Barmada M.M., Finegold D.N.;
     "Hereditary lymphedema: evidence for linkage and genetic
RТ
     heterogeneity.";
RT
     Hum. Mol. Genet. 7:2073-2078(1998).
RL
RN
RР
     INVOLVEMENT IN HEREDITARY LYMPHEDEMA I, AND CHARACTERIZATION OF
     VARIANT HEREDITARY LYMPHEDEMA I ARG-1035.
RP
RX
     PubMed=10856194;
     Irrthum A., Karkkainen M.J., Devriendt K., Alitalo K., Vikkula M.;
RA
     "Congenital hereditary lymphedema caused by a mutation that
RT
     inactivates VEGFR3 tyrosine kinase.";
RT
     Am. J. Hum. Genet. 67:295-301(2000).
RL
RN
     [9]
RΡ
     VARIANTS HEREDITARY LYMPHEDEMA I ARG-857; PRO-1041; PRO-1044 AND
RP
     LEU-1114, VARIANT SER-641, AND CHARACTERIZATION OF VARIANTS.
     PubMed=10835628; DOI=10.1038/75997;
RX
RA
     Karkkainen M.J., Ferrell R.E., Lawrence E.C., Kimak M.A.,
     Levinson K.L., McTigue M.A., Alitalo K., Finegold D.N.;
RA
     "Missense mutations interfere with VEGFR-3 signalling in primary
RT
RT
     lymphoedema.";
RL
     Nat. Genet. 25:153-159(2000).
RN
RP
     VARIANTS JUVENILE HEMANGIOMA SER-954 AND SER-1137, AND VARIANTS
RP
     ALA-494; GLN-890 AND HIS-1146.
     PubMed=11807987; DOI=10.1002/gcc.10028;
RX
     Walter J.W., North P.E., Waner M., Mizeracki A., Blei F.,
     Walker J.W.T., Reinisch J.F., Marchuk D.A.;
RA
RT
     "Somatic mutation of vascular endothelial growth factor receptors in
RT
     juvenile hemangioma.";
RL
     Genes Chromosomes Cancer 33:295-303(2002).
CC
     -!- FUNCTION: Receptor for VEGFC. Has a tyrosine-protein kinase
CC
         activity.
     -!- CATALYTIC ACTIVITY: ATP + a protein tyrosine = ADP + protein
CC
CC
         tyrosine phosphate.
CC
     -!- SUBCELLULAR LOCATION: Type I membrane protein.
CC
     -!- TISSUE SPECIFICITY: Placenta, lung, heart, and kidney, does not
CC
         seem to be expressed in pancreas and brain.
CC
     -!- DISEASE: Defects in FLT4 are the cause of hereditary lymphedema I
CC
         [MIM:153100]; also known as Nonne-Milroy lymphedema or Milroy
         disease. Hereditary lymphedema is a chronic disabling condition
CC
CC
         which results in swelling of the extremities due to altered
CC
         lymphatic flow. Patients with lymphedema suffer from recurrent
         local infections and physical impairment. Hereditary lymphedema I
CC
CC
         shows autosomal dominant inheritance and is characterized by onset
         usually at birth.
```

```
CC
     -!- DISEASE: Defects in FLT4 are found in juvenile hemangioma.
         Juvenile hemangiomas are the most common tumors of infancy,
CC
         occurring as many as 10/% of all births. These benign vascular
CC
CC
         lesions enlarge rapidly during the first year of life by
CC
         hyperplasia of endothelial cells and attendant pericytes, and then
CC
         spontaneously involute over a period of years, leaving loose
CC
         fibrofatty tissue.
CC
     -!- SIMILARITY: Belongs to the Tyr protein kinase family. CSF-1/PDGF
CC
         receptor subfamily.
CC
     -!- SIMILARITY: Contains 7 immunoglobulin-like C2-type domains.
CC
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     entities requires a license agreement (See http://www.isb-sib.ch/announce/
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     or send an email to license@isb-sib.ch).
CC
     EMBL; X69878; CAA49505.1; -.
DR
     EMBL; X68203; CAA48290.1; ALT_INIT.
     EMBL; U43143; AAA85215.1; -.
DR
DR
     PIR; A48999; A48999.
DR
     HSSP; P35968; 1VR2.
DR
     Genew; HGNC: 3767; FLT4.
DR
     MIM; 136352; -.
DR
     MIM; 153100; -.
     GO; GO:0005887; C:integral to plasma membrane; TAS.
DR
DR
     GO; GO:0005021; F:vascular endothelial growth factor receptor. . .; TAS.
     GO; GO:0007169; P:transmembrane receptor protein tyrosine kin. . .; TAS.
DR
     InterPro; IPR007110; Ig-like.
DR
DR
     InterPro; IPR003598; Ig_c2.
DR
     InterPro; IPR011009; Kinase_like.
DR
     InterPro; IPR000719; Prot kinase.
DR
     InterPro; IPR001824; RecepttyrkinsIII.
DR
     InterPro; IPR001245; Tyr_pkinase.
     InterPro; IPR008266; Tyr_pkinase_AS.
DR
DR
     InterPro; IPR009134; VEGFR.
DR
     InterPro; IPR009137; VEGFR3.
     Pfam; PF00047; ig; 6.
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DR
DR
     PRINTS; PR01832; VEGFRECEPTOR.
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     PRINTS; PR01835; VEGFRECEPTR3.
DR
     ProDom; PD000001; Prot kinase; 2.
DR
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DR
     SMART; SM00219; TyrKc; 1.
     PROSITE; PS50835; IG LIKE; 6.
DR
     PROSITE; PS00107; PROTEIN_KINASE_ATP; 1.
DR
     PROSITE; PS50011; PROTEIN KINASE DOM; 1.
DŔ
     PROSITE; PS00109; PROTEIN_KINASE_TYR; 1.
DR
DR
     PROSITE; PS00240; RECEPTOR_TYR_KIN_III; 1.
     ATP-binding; Direct protein sequencing; Disease mutation;
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KW
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KW
     Receptor; Repeat; Signal; Transferase; Transmembrane;
KW
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     SIGNAL
FT
FT
     CHAIN
                  25
                       1298
                                  Vascular endothelial growth factor
FT
                                   receptor 3.
                                                                          ...
FT
     DOMAIN
                  25
                        775
                                  Extracellular (Potential).
FT
     TRANSMEM
                 776
                        797
                                   Potential.
FT
     DOMAIN
                 798
                       1298
                                  Cytoplasmic (Potential).
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     DOMATN
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FT
     DOMAIN
                 151
                        213
                                  Ig-like C2-type 2.
     DOMAIN
FT
                 219
                        326
                                  Ig-like C2-type 3.
FT
     DOMAIN
                 331
                        415
                                  Ig-like C2-type 4.
FT
     DOMAIN
                                  Ig-like C2-type 5.
                 422
                        552
FT
     DOMAIN
                 555
                        671
                                  Ig-like C2-type 6.
FΤ
     DOMAIN
                 678
                        764
                                  Ig-like C2-type 7.
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FT
                 845
                       1173
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                                  ATP (By similarity).
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                 879
                        879
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                    111
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    DISULFID
              252
                             Potential.
FT
                    310
    DISULFID
                             Potential.
              445
FT
                    534
    DISULFID
              578
                             Potential.
FT
                    653
    DISULFID
              699
                    751
                             Potential.
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FT
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               33
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    CARBOHYD
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                             N-linked (GlcNAc. . .) (Potential) .
FT
                    104
                             N-linked (GlcNAc. . .) (Potential).
FT
    CARBOHYD
              166
                    166
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N-linked (GlcNAc. . .) (Potential) .
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              251
                    251
    CARBOHYD
FT
              299
                    299
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                             N-linked (GlcNAc. . .) (Potential).
FT
              411
                    411
                             N-linked (GlcNAc. . .) (Potential).
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FT
              515
                    515
    CARBOHYD
              527
                             N-linked (GlcNAc. . .) (Potential).
FT
                    527
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                             N-linked (GlcNAc. . .) (Potential).
              594
FT
                    594
FT
    CARBOHYD
              683
                    683
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FT
    CARBOHYD
              690
                    690
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                             N-linked (GlcNAc. . .) (Potential).
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    CARBOHYD
              758
                    758
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FT
    MOD RES
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                    1068
                             similarity).
FT
FT
    VARIANT
              494
                    494
                             /FTId=VAR 018407.
FT
FT
    VARIANT
              641
                    641
                             P -> S (does not affect kinase activity).
                             /FTId=VAR_018408.
FT
FT
    VARIANT
              857
                    857
                             G -> R (in hereditary lymphedema I; loss
FT
                             of kinase activity).
  Query Match
                      95.0%; Score 6852; DB 1; Length 1298;
 Best Local Similarity
                     100.0%; Pred. No. 0;
                            0; Mismatches
 Matches 1297: Conservative
                                              Indels
          1 MQRGAALCLRLWLCLGLLDGLVSGYSMTPPTLNITEESHVIDTGDSLSISCRGQHPLEWA 60
Qу
            Db
          1 MQRGAALCLRLWLCLGLLDGLVSGYSMTPPTLNITEESHVIDTGDSLSISCRGQHPLEWA 60
         61 WPGAQEAPATGDKDSEDTGVVRDCEGTDARPYCKVLLLHEVHANDTGSYVCYYKYIKARI 120
Qу
            Dh
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        121 EGTTAASSYVFVRDFEQPFINKPDTLLVNRKDAMWVPCLVSIPGLNVTLRSQSSVLWPDG 180
Qу
            Db
        121 EGTTAASSYVFVRDFEOPFINKPDTLLVNRKDAMWVPCLVSIPGLNVTLRSOSSVLWPDG 180
        181 QEVVWDDRRGMLVSTPLLHDALYLQCETTWGDQDFLSNPFLVHITGNELYDIQLLPRKSL 240
Oν
            Db
           QEVVWDDRRGMLVSTPLLHDALYLQCETTWGDQDFLSNPFLVHITGNELYDIQLLPRKSL 240
        241 ELLVGEKLVLNCTVWAEFNSGVTFDWDYPGKQAERGKWVPERRSQQTHTELSSILTIHNV 300
Qy
            Db
           ELLVGEKLVLNCTVWAEFNSGVTFDWDYPGKQAERGKWVPERRSQCTHTELSSILTIHNV 300
        301 SQHDLGSYVCKANNGIQRFRESTEVIVHENPFISVEWLKGPILEATAGDELVKLPVKLAA 360
Qу
            Db
           SOHDLGSYVCKANNGIORFRESTEVIVHENPFISVEWLKGPILEATAGDELVKLPVKLAA 360
Qу
        361 YPPPEFQWYKDGKALSGRHSPHALVLKEVTEASTGTYTLALWNSAAGLRRNISLELVVNV 420
             Db
        361 YPPPEFOWYKDGKALSGRHSPHALVLKEVTEASTGTYTLALWNSAAGLRRNISLELVVNV 420
           PPQIHEKEASSPSIYSRHSRQALTCTAYGVPLPLSIQWHWRPWTPCKMFAQRSLRRRQQQ 480
Qу
            Db
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Qу
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        481 DLMPQCRDWRAVTTQDAVNPIESLDTWTEFVEGKNKTVSKLVIQNANVSAMYKCVVSNKV 540
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Ον
           661 HDKHCHKKYLSVQALEAPRLTQNLTDLLVNVSDSLEMQCLVAGAHAPSIVWYKDERLLEE 720
        721 KSGVDLADSNQKLSIQRVREEDAGRYLCSVCNAKGCVNSSASVAVEGSEDKGSMEIVILV 780
Qy .
           Db
          KSGVDLADSNOKLSIORVREEDAGRYLCSVCNAKGCVNSSASVAVEGSEDKGSMEIVILV 780
        781 GTGVIAVFFWVLLLLIFCNMRRPAHADIKTGYLSIIMDPGEVPLEEQCEYLSYDASQWEF 840
Qу
           781 GTGVIAVFFWVLLLLIFCNMRRPAHADIKTGYLSIIMDPGEVPLEEQCEYLSYDASQWEF 840
Db
        841 PRERLHLGRVLGYGAFGKVVEASAFGIHKGSSCDTVAVKMLKEGATASEHRALMSELKIL 900
Qy
           PRETLHLGRVLGYGAFGKVVEASAFGIHKGSSCDTVAVKMLKEGATASEHRALMSELKIL 900
Db
        901 IHIGNHLNVVNLLGACTKPQGPLMVIVEFCKYGNLSNFLRAKRDAFSPCAEKSPEQRGRF 960
Qу
           Db
        901 IHIGNHLNVVNLLGACTKPQGPLMVIVEFCKYGNLSNFLRAKRDAFSPCAEKSPEQRGRF 960
Qу
          RAMVELARLDRRRPGSSDRVLFARFSKTEGGARRASPDQEAEDLWLSPLTMEDLVCYSFQ 1020
           RAMVELARLDRRRPGSSDRVLFARFSKTEGGARRASPDQEAEDLWLSPLTMEDLVCYSFQ 1020
       1021 VARGMEFLASRKCIHRDLAARNILLSESDVVKICDFGLARDIYKDPDYVRKGSARLPLKW 1080
Qy
           Db
       1021 VARGMEFLASRKCIHRDLAARNILLSESDVVKICDFGLARDIYKDPDYVRKGSARLPLKW 1080
       1081 MAPESIFDKVYTTQSDVWSFGVLLWEIFSLGASPYPGVQINEEFCQRLRDGTRMRAPELA 1140
Qу
           1081 MAPESIFDKVYTTQSDVWSFGVLLWEIFSLGASPYPGVQINEEFCQRLRDGTRMRAPELA 1140
Db
       1141 TPAIRRIMLNCWSGDPKARPAFSELVEILGDLLOGRGLOEEEVCMAPRSSOSSEEGSFS 1200
Oν
           Db
       1141 TPAIRRIMLNCWSGDPKARPAFSELVEILGDLLQGRGLQEEEEVCMAPRSSQSSEEGSFS 1200
       1201 QVSTMALHIAQADAEDSPPSLQRHSLAARYYNWVSFPGCLARGAETRGSSRMKTFEEFPM 1260
Qγ
           1201 QVSTMALHIAQADAEDSPPSLQRHSLAARYYNWVSFPGCLARGAETRGSSRMKTFEEFPM 1260
Db
       1261 TPTTYKGSVDNQTDSGMVLASEEFEQIESRHRQESGF 1297
Qy
           Dh
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    P35968; O60723; Q14178;
AC
DT
   01-JUN-1994 (Rel. 29, Created)
   16-OCT-2001 (Rel. 40, Last sequence update)
DT
DT
   25-OCT-2004 (Rel. 45, Last annotation update)
   Vascular endothelial growth factor receptor 2 precursor (EC 2.7.1.112)
    (VEGFR-2) (Kinase insert domain receptor) (Protein-tyrosine kinase
DE
   receptor Flk-1).
DE
   Name=KDR; Synonyms=FLK1;
GN
os
   Homo sapiens (Human).
OC
   Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;
OC
   Mammalia; Eutheria; Primates; Catarrhini; Hominidae; Homo.
ox
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RN
    [1]
   SEQUENCE FROM N.A.
RP
   Yin L.Y., Wu Y., Patterson C.;
RA
RT
    "Full length human KDR/flk-1 sequence.";
   Submitted (DEC-1997) to the EMBL/GenBank/DDBJ databases.
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RN

[2]

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RP
     SEQUENCE FROM N.A.
RC
     TISSUE=Umbilical vein;
RA
     Yu Y., Whitney R.G., Sato J.D.;
     "Coding region for human VEGF receptor KDR (VEGFR-2).";
RT
     Submitted (MAY-1998) to the EMBL/GenBank/DDBJ databases.
RL
RN
RP
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RC
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     MEDLINE=92019839; PubMed=1656371;
RA
     Terman B.I., Carrion M.E., Kovacs E., Rasmussen B.A., Eddy R.L.,
RA
     Shows T.B.;
RT
     "Identification of a new endothelial cell growth factor receptor
RT
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RT.
     Oncogene 6:1677-1683(1991).
RN
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RΡ
RX
     MEDLINE=96032749; PubMed=7559454; DOI=10.1074/jbc.270.39.23111;
RA
     Patterson C., Perrella M.A., Hsieh C.M., Yoshizumi M., Lee M.E.,
RA
     Harber E.;
     "Cloning and functional analysis of the promoter for KDR/flk-1, a
RT
     receptor for vascular endothelial growth factor.";
RL
     J. Biol. Chem. 270:23111-23118(1995).
RN
     [5]
RP
     FUNCTION.
RX
     MEDLINE=93038639; PubMed=1417831;
RA
     Terman B.I., Dougher-Vermazen M., Carrion M.E., Dimitrov D.,
     Armellino D.C., Gospodarowicz D., Boehlen P.;
RA
RT
     "Identification of the KDR tyrosine kinase as a receptor for vascular
RT
     endothelial cell growth factor.";
     Biochem. Biophys. Res. Commun. 187:1579-1586(1992).
RL
     -!- FUNCTION: Receptor for VEGF or VEGFC. Has a tyrosine-protein
CC
CC
         kinase activity. The VEGF-kinase ligand/receptor signaling system
CC
         plays a key role in vascular development and regulation of
CC
         vascular permeability.
CC
     -!- CATALYTIC ACTIVITY: ATP + a protein tyrosine = ADP + protein
CC
         tyrosine phosphate.
     -!- SUBCELLULAR LOCATION: Type I membrane protein.
CC
CC
     -!- SIMILARITY: Belongs to the Tyr protein kinase family. CSF-1/PDGF
CC
        receptor subfamily.
CC
     -!- SIMILARITY: Contains 7 immunoglobulin-like C2-type domains.
CC
     ------
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     or send an email to license@isb-sib.ch).
CC
DR
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DR
     EMBL; AF063658; AAC16450.1; -.
     EMBL; X61656; CAA43837.1; -.
DR
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DR
DR
     PDB; 1VR2; X-ray; A=806-1171.
DR
    Genew; HGNC:6307; KDR.
DR
    MIM; 191306; -.
     GO; GO:0005887; C:integral to plasma membrane; TAS.
DR
    GO; GO:0005021; F:vascular endothelial growth factor receptor. . .; TAS.
     GO; GO:0007169; P:transmembrane receptor protein tyrosine kin. . .; TAS.
DR
    InterPro; IPR007110; Ig-like.
DR
DR
     InterPro; IPR003598; Ig_c2.
DR
     InterPro; IPR011009; Kinase_like.
DR
     InterPro; IPR000719; Prot_kinase.
     InterPro; IPR001824; RecepttyrkinsIII.
DR
     InterPro; IPR001245; Tyr_pkinase.
DR
     InterPro; IPR008266; Tyr_pkinase_AS.
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DR
     SMART; SM00408; IGc2; 2.
DR
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DR
DR
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     PROSITE; PS50011; PROTEIN_KINASE_DOM; 1.
DR
DR
     PROSITE; PS00109; PROTEIN_KINASE_TYR; 1.
DR
     PROSITE; PS00240; RECEPTOR_TYR_KIN_III; 1.
KW
     3D-structure; Angiogenesis; ATP-binding; Glycoprotein;
KW
     Immunoglobulin domain; Phosphorylation; Receptor; Repeat; Signal;
     Transferase; Transmembrane; Tyrosine-protein kinase.
KW
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FT
     SIGNAL
                    1
                          19
                                     Vascular endothelial growth factor
     CHAIN
FT
                   20
                         1356
FT
                                     receptor 2.
                                     Extracellular (Potential).
FT
     DOMAIN
                   20
                          764
FT
     TRANSMEM
                  765
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                                     Potential.
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     DOMAIN
                  790
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FT
                   46
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FT
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                                     Ig-like C2-type 3.
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                                     Ig-like C2-type 5.
FΤ
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                                     Ig-like C2-type 7.
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N-linked (GlcNAc. . .) (Potential) .
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FT
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FT
                          96
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                                     N-linked (GlcNAc. . .) (Potential).
FT
                  143
                          143
FT
     CARBOHYD
                  158
                          158
                                    N-linked (GlcNAc. . .) (Potential).
FΤ
     CARBOHYD
                  245
                          245
                                    N-linked (GlcNAc. . .) (Potential).
                                    N-linked (GlcNAc. . .) (Potential).
FT
     CARBOHYD
                  318
                          318
                                    N-linked (GlcNAc. . .) (Potential).
FT
     CARBOHYD
                  374
                          374
                                    N-linked (GlcNAc. . .) (Potential).
N-linked (GlcNAc. . .) (Potential).
FT
     CARBOHYD
                  395
                          395
     CARBOHYD
FT
                  511
                          511
     CARBOHYD
                                    N-linked (GlcNAc. . .) (Potential).
                  523
                          523
FТ
     CARBOHYD
                  580
                          580
                                    N-linked (GlcNAc. . .) (Potential).
FT
     CARBOHYD
                  613
                          613
                                    N-linked (GlcNAc. . .) (Potential).
     CARBOHYD
                                    N-linked (GlcNAc. . .) (Potential).
FT
                  619
                         619
FT
     CARBOHYD
                  631
                          631
                                    N-linked (GlcNAc. . .) (Potential).
FΤ
     CARBOHYD
                  675
                          675
                                    N-linked (GlcNAc. . .) (Potential).
                                    N-linked (GlcNAc. . .) (Potential).
FT
     CARBOHYD
                  704
                          704
                                    N-linked (GlcNAc. . .) (Potential) .
FT
     CARBOHYD
                  721
                          721
FT
     MOD_RES
                 1059
                        1059
                                    Phosphotyrosine (by autocatalysis) (By
                                    similarity).
FT
     CONFLICT
FT
                                    Q \rightarrow E (in Ref. 2).
FT
     CONFLICT
                  772
                         772
                                    A \rightarrow T \text{ (in Ref. 3)}.
FT
     CONFLICT
                  787
                                    R \rightarrow G \text{ (in Ref. 3)}.
                          787
FT
     CONFLICT
                  835
                          835
                                    K \rightarrow N \text{ (in Ref. 3)}.
FT
     CONFLICT
                  848
                         848
                                    V \rightarrow E \text{ (in Ref. 3)}.
FT
     CONFLICT
                 1347
                        1347
                                    S \rightarrow T (in Ref. 3).
                 1356 AA; 151526 MW; 59E7C44B05CFEBB3 CRC64;
     SEQUENCE
  Query Match 38.3%; Score 2762; DB 1; Length 1356; Best Local Similarity 44.1%; Pred. No. 8.5e-172;
  Matches 608; Conservative 203; Mismatches 480; Indels
                                                                     88; Gaps
Qy
             1 MQRGAALCLRLWLCLGLLDGLVS--GYSMTPPTLNITEESHVIDTGDSLSISCRGQHPLE 58
                                   11 | 1 | 1 | 1 | 1 | 1
             1 MOSKVLLAVALWLCVETRAASVGLPSVSLDLPRLSIOKDILTIKANTTLOITCRGORDLD 60
Db
              WAWPGAQEAPATGDKDSEDTGVVRDC-EGTDARPYCKVLLLHEVHANDTGSYVCYYKYIK 117
Qу
                                -SEQRVEVTECSDGL----FCKTLTIPKVIGNDTGAYKCFYR--- 106
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Qу	118	ARIEGTTAASSYVFVRDFEQPFINKPDTLLVNRKDAMWVPCLVSIPGLNVTL	169
Db	107	ETDLASVIYVYVQDYRSPFIASVSDQHGVVYITENKNKTVVIPCLGSISNLNVSLCA	163
Qy	170	RSQSSVLWPDGQEVVWDDRRGMLVSTPLLHDALYLQCETTWGDQDFLSNPFLVHITGNEL	229
Db	164	RYPEKRFVPDGNRISWDSKKGFTIPSYMISYAGMVFCEAKINDESYQSIMYIVVVVGYRI	223
Qy		YDIQLLPRKSLELLVGEKLVLNCTVWAEFNSGVTFDWDYPGKQAERGKWVPERRSQQTHT :	
Db	224	YDVVLSPSHGIELSVGEKLVLNCTARTELNVGIDFNWEYPSSKHQHKKLVNRDLKTQSGS	
Qy		ELSSILTIHNVSQHDLGSYVCKANNGIQRFRESTEVIVHENPFISVEWLKGPILEAT : :: :: : :: ::	
Db		EMKKFLSTLTIDGVTRSDQGLYTCAASSGLMTKKNSTFVRVHEKPFVAFGSGMESLVEAT	
Qy Db		AGDELVKLPVKLAAYPPPEFQWYKDGKALSGRHSPHALVLKEVTEASTGTYTLALWN	,
Qy		SAAGLRRNISLELVVNVPPQIHEKEASSP-SIYSRHSRQALTCTAYGVPLPLSIQWHWRP	
Db		: ::: : ' : : : PISKEKQSHVVSLVVYVPPQIGEKSLISPVDSYQYGTTQTLTCTVYAIPPPHHIHWYWQL	
Qy		WTPCKMFAQRSLRRRQQQDLMPQCRDWRAVTTQDAVNPIESLDTWTEFVEGKNKTVSKLV	
Db		: : : : : :	
Qy		IQNANVSAMYKCVVSNKVGQDERLIYFYVTTIPDGFTIESKPSEELLEGQPVLLSCQADS	
Db		: : : : : : :	575
Qy	583	YKYEHLRWYRLNLSTLHDAHGNPLLLDCKNVHLFATPLAASLEEVAPGARHATLSL	638
Db	576	: : :	627
Qy	639	SIPRVAPEHEGHYVCEVQDRRSHDKHCHKKYLSVQALEAPRLTQNLTDLLVNVSDSLEMQ : :: : : : : : : :	698
Db	628	: : : : : :	687
Qy	699	CLVAGAHAPSIVWYKDERLLEEKSGVDLADSNQKLSIQRVREEDAGRYLCSVCNAKGCVN	758
Db	688	CTASGNPPPQIMWFKDNETLVEDSGIVLKDGNRNLTIRRVRKEDEGLYTCQACSVLGCAK	747
Qy	759	SSASVAVEGSEDKGSMEIVILVGTGVIAVFFWVLLLLIFCNMRRPAHADIKTGYLSIIMD : ::: :: : : :: :: ::	818
Db		VEAFFIIEGAQEKTNLEIIILVGTAVIAMFFWLLLVIILRTVKRANGGELKTGYLSIVMD	
Qу		PGEVPLEEQCEYLSYDASQWEFPRERLHLGRVLGYGAFGKVVEASAFGIHKGSSCDTVAV : : : : : : :	
Db	•	PDELPLDEHCERLPYDASKWEFPRDRLKLGKPLGRGAFGQVIEADAFGIDKTATCRTVAV	
Qy		KMLKEGATASEHRALMSELKILIHIGNHLNVVNLLGACTKPQGPLMVIVEFCKYGNLSNF	
Db		KMLKEGATHSEHRALMSELKILIHIGHHLNVVNLLGACTKPGGPLMVIVEFCKFGNLSTY	
Qy Db		LRAKRDAFSPCAEKSPEQRGRFRAMVELARLDRRRPGSSDRVLFARFSKTEGGARRA : : : :: : LRSKRNEFVPYKTKGARFRQGKDYVGAIPVDLKRRLDSITSSQSSASSGFVEEK	
Qy		SPDQEA-EDLWLSPLTMEDLVCYSFQVARGMEFLASRKCIHRDLAARNILLSESDV	
Db		:: : : : : : : : :	
	1051	VKICDFGLARDIYKDPDYVRKGSARLPLKWMAPESIFDKVYTTQSDVWSFGVLLWEIFSL	
Db			1101
Qу		GASPYPGVQINEEFCQRLRDGTRMRAPELATPAIRRIMLNCWSGDPKARPAFSELVEILG	1170
Db		: : : :	1161

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1171 DLLQGRGLQEEEEVCMAPRS-SQSSEEGSFSQVSTMALHIAQADAEDSPPSLQRHSLAAR 1229
Qy
              DЪ
         1162 NLLQANAQODGKDYIVLPISETLSMEEDSGLSLPTSPVSCMEEEEVCDP------KFH 1213
         1230 YYNWVSFPGCLARGAETRGSSRMKTFEEFPM-TPTTYKGSVDNQTDSGMVLASEEFEQIE 1288
Qy
                                    :||||: |: |
              1 1
                                                       DЬ
         1214 YDNTAGISQYLQNSKRKSRPVSVKTFEDIPLEEPEVKVIPDDNQTDSGMVLASEELKTLE 1273
         1289 SRHRQESGFSCKGPGQNVAVTRAHPDSQGRRRRPERGARGGQVFYNSEYGELSEPSEED 1347
Qу
                        | : :| |:| :
                                                   1 :
Db
         1274 DRTKLSPSFGGMVPSK----SRESVASEGSNQ----TSGYQSGYHSDDTDTTVYSSEE 1323
RESULT 12
VGR2 MOUSE
     VGR2_MOUSE
                                   PRT; 1367 AA.
ID
                    STANDARD:
AC
     P35918;
DT
     01-JUN-1994 (Rel. 29, Created)
     01-JUN-1994 (Rel. 29, Last sequence update) 05-JUL-2004 (Rel. 44, Last annotation update)
DT
DT
     Vascular endothelial growth factor receptor 2 precursor (EC 2.7.1.112)
     (VEGFR-2) (Protein-tyrosine kinase receptor flk-1) (Fetal liver kinase
DE
DΕ
     1) (Kinase NYK).
     Name=Kdr; Synonyms=Flk-1, Flk1;
GN
os
     Mus musculus (Mouse).
OC
     Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;
OC
     Mammalia; Eutheria; Rodentia; Sciurognathi; Muridae; Murinae; Mus.
ox
     NCBI_TaxID=10090;
RN
     [1]
     SEQUENCE FROM N.A.
RP
     STRAIN=BALB/c; TISSUE=Embryo;
RC
     MEDLINE=93208880; PubMed=7681362; DOI=10.1016/0092-8674(93)90573-9;
RA
     Millauer B., Wizigmann-Voos S., Schnurch H., Martinez R.,
RA
     Mueller N.P.H., Risau W., Ullrich A.;
     "High affinity VEGF binding and developmental expression suggest Flk-1
RT
RT
     as a major regulator of vasculogenesis and angiogenesis.";
RL
     Cell 72:835-846(1993).
RN
     [2]
     SEQUENCE FROM N.A.
RP
RC
     STRAIN=C3H/He; TISSUE=Fetal liver;
RX
     MEDLINE=92020984; PubMed=1717995;
RA
     Mathews W., Jordan C.T., Gavin M., Jenkins N.A., Copeland N.G.,
RA
     Lemishcka I.R.;
RT
     "A receptor tyrosine kinase cDNA isolated from a population of
RT
     enriched primitive hematopoietic cells and exhibiting close genetic
     linkage to c-kit.";
     Proc. Natl. Acad. Sci. U.S.A. 88:9026-9030(1991).
RL
RN
     [3]
RP
     SEQUENCE FROM N.A.
RX
     MEDLINE=93141255; PubMed=8423988;
RA
     Oelrichs R.B., Reid H.H., Bernard O., Ziemiecki A., Wilks A.F.;
RT
     "NYK/FLK-1: a putative receptor protein tyrosine kinase isolated from
RT
     E10 embryonic neuroepithelium is expressed in endothelial cells of the
PТ
     developing embryo.";
RL
     Oncogene 8:11-18(1993).
RN
     SEQUENCE OF 1-15 FROM N.A.
RP
     MEDLINE=96032749; PubMed=7559454; DOI=10.1074/jbc.270.39.23111;
RX
RA
     Patterson C., Perrella M.A., Hsieh C.-M., Yoshizumi M., Lee M.-E.,
RA
     Harber E.;
RT
     "Cloning and functional analysis of the promoter for KDR/flk-1, a
     receptor for vascular endothelial growth factor.";
RT
RI.
     J. Biol. Chem. 270:23111-23118(1995).
RN
     [5]
RP
    FUNCTION.
RX
    MEDLINE=93361481; PubMed=8356051;
     Quinn T.P., Peters K.G., de Vries C., Ferrara N., Williams L.T.;
RA
     "Fetal liver kinase 1 is a receptor for vascular endothelial growth
```

```
factor and is selectively expressed in vascular endothelium.";
RT
RL
     Proc. Natl. Acad. Sci. U.S.A. 90:7533-7537(1993).
CC
     -!- FUNCTION: Receptor for VEGF or VEGFC. Has a tyrosine-protein
CC
         kinase activity. The VEGF-kinase ligand/receptor signaling system
CC
         plays a key role in vascular development and regulation of
CC
         vascular permeability.
CC
     -!- CATALYTIC ACTIVITY: ATP + a protein tyrosine = ADP + protein
CC
         tyrosine phosphate.
     -!- SUBCELLULAR LOCATION: Type I membrane protein.
CC
CC
     -!- TISSUE SPECIFICITY: Expressed at high levels in adult heart, lung,
CC
         kidney, brain and skeletal muscle, but is also expressed at lower
CC
         levels in most other adult tissues.
CC
     -!- SIMILARITY: Belongs to the Tyr protein kinase family. CSF-1/PDGF
CC
         receptor subfamily.
CC
     -!- SIMILARITY: Contains 7 immunoglobulin-like C2-type domains.
CC
     ______
CC
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CC
     between the Swiss Institute of Bioinformatics and the EMBL outstation -
     the European Bioinformatics Institute. There are no restrictions on its
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     or send an email to license@isb-sib.ch).
CC
CC
     _____
DR
     EMBL; X70842; CAA50192.1; -.
DR
     EMBL; X59397; CAA42040.1; -.
DR
     EMBL; S53103; AAB25043.1; -.
DR
     EMBL; X89777; CAA61917.1; -.
DR
     PIR; A41228; A41228.
     HSSP; P35968; 1VR2.
DR
     MGD; MGI:96683; Kdr.
DR
DR
     GO; GO:0045165; P:cell fate commitment; IMP.
DR
     GO; GO:0045446; P:endothelial cell differentiation; IDA.
DR
     InterPro; IPR007110; Ig-like.
DR
     InterPro; IPR003598; Ig_c2.
DR
     InterPro; IPR011009; Kinase_like.
     InterPro; IPR000719; Prot_kinase.
DR
     InterPro; IPR001824; RecepttyrkinsIII.
DR
     InterPro; IPR001245; Tyr_pkinase.
     InterPro; IPR008266; Tyr pkinase AS.
DR
     InterPro; IPR009134; VEGFR.
DR
DR
     InterPro; IPR009136; VEGFR2.
     Pfam; PF00047; ig; 6.
DR
     Pfam; PF00069; Pkinase; 1.
DR
DR
     PRINTS; PR01832; VEGFRECEPTOR.
DR
     PRINTS; PR01834; VEGFRECEPTR2.
     ProDom; PD000001; Prot_kinase; 2.
DR
     SMART; SM00408; IGc2; 1.
DR
DR
     SMART; SM00219; TyrKc; 1.
     PROSITE; PS50835; IG_LIKE; 5.
DR
DR
     PROSITE; PS00107; PROTEIN KINASE ATP; 1.
DR
     PROSITE; PS50011; PROTEIN KINASE DOM; 1.
DR
     PROSITE; PS00109; PROTEIN_KINASE_TYR; 1.
DR
     PROSITE; PS00240; RECEPTOR_TYR_KIN_III; 1.
KW
     Angiogenesis; ATP-binding; Glycoprotein; Immunoglobulin domain;
     Phosphorylation; Receptor; Repeat; Signal; Transferase; Transmembrane;
KW
     Tyrosine-protein kinase.
FT
     SIGNAL
                  1
                        19
                                  Potential.
FT
     CHAIN
                 20
                      1367
                                  Vascular endothelial growth factor
FT
                                  receptor 2.
FT
     DOMAIN
                 20
                        762
                                  Extracellular (Potential).
                 763
FT
     TRANSMEM
                        784
                                  Potential.
FT
     DOMAIN
                 785
                       1367
                                  Cytoplasmic (Potential).
FT
     DOMAIN
                 46
                                  Ig-like C2-type 1.
                       111
FT
     DOMAIN
                 143
                        209
                                  Ig-like C2-type 2.
                                  Ig-like C2-type 3.
FT
     DOMAIN
                 226
                        325
     DOMAIN
FT
                 330
                        416
                                  Ig-like C2-type 4.
                                  Ig-like C2-type 5.
FT
     DOMAIN
                 423
                        542
    DOMAIN
FT
                 549
                        656
                                  Ig-like C2-type 6.
     DOMAIN
                                  Ig-like C2-type 7.
FT
                 665
                        751
    DOMAIN
                 832
                      1160
                                 Protein kinase.
```

```
NP_BIND
                               ATP (By similarity).
FT
               838
                     846
                               ATP (By similarity).
FT
    BINDING
               866
                      866
FT
    ACT SITE
              1026
                     1026
                               By similarity.
FΤ
    CARBOHYD
                46
                               N-linked (GlcNAc. . .) (Potential).
                      46
    CARBOHYD
                               N-linked (GlcNAc. . .) (Potential).
FT
                98
                      98
    CARBOHYD
               145
                               N-linked (GlcNAc. . .) (Potential).
FT
                     145
    CARBOHYD
               160
                               N-linked (GlcNAc. . .) (Potential).
FT
                     160
FT
    CARBOHYD
               247
                      247
                               N-linked (GlcNAc. . .) (Potential).
    CARBOHYD
                               N-linked (GlcNAc. . .) (Potential).
FT
               320
                      320
                               N-linked (GlcNAc. . .) (Potential).
FT
    CARBOHYD
               376
                      376
                               N-linked (GlcNAc. . .) (Potential).
N-linked (GlcNAc. . .) (Potential).
FT
    CARBOHYD
               397
                      397
FT
    CARBOHYD
               509
                      509
    CARBOHYD
               521
                      521
                               N-linked (GlcNAc. . .) (Potential).
    CARBOHYD
                               N-linked (GlcNAc. . .) (Potential).
               578
                     578
FΤ
    CARBOHYD
                               N-linked (GlcNAc. . .) (Potential).
FT
               611
                      611
                               N-linked (GlcNAc. . .) (Potential).
    CARBOHYD
FT
               617
                     617
FT
    CARBOHYD
               629
                      629
                               N-linked (GlcNAc. . .) (Potential).
FT
    CARBOHYD
               673
                      673
                               N-linked (GlcNAc. . .) (Potential).
                               N-linked (GlcNAc. . .) (Potential).
FT
    CARBOHYD
               702
                      702
                               N-linked (GlcNAc. . .) (Potential).
FT
    CARBOHYD
               719
                      719
    MOD_RES
                     1057
                               Phosphotyrosine (by autocatalysis) (By
FT
              1057
                               similarity).
    CONFLICT
                25
                      25
                               P \rightarrow T (in Ref. 1).
FT
                               G -> D (in Ref. 3).
FT
    CONFLICT
               679
                      679
    CONFLICT
               783
                     784
                               LV -> VL (in Ref. 1).
FT
FT
    CONFLICT
               917
                     917
                               S -> C (in Ref. 1).
    CONFLICT
                               QLTSCLNGSGPVPAPPPTPGNHERGAA -> RSPPV (in
FT
              1341
                     1367
FΤ
                               Ref. 3).
    SEQUENCE
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 Query Match 37.0%; Score 2669; DB 1; Length 1367; Best Local Similarity 43.0%; Pred. No. 1.1e-165;
 Matches 588; Conservative 205; Mismatches 485; Indels
                                                          90; Gaps
           1 MQRGAALCLRLWLCLGLLDGLVS--GYSMTPPTLNITEESHVIDTGDSLSISCRGQHPLE 58
Oν
             1 MESKALLAVALWFCVETRAASVGLPGDFLHPPKLSTQKDILTILANTTLQITCRGQRDLD 60
Db
          59 WAWPGAQEAPATGDKDSEDTGVVRDCEGTDARPYCKVLLLHEVHANDTGSYVCYYKYIKA 118
Qу
                    1 11 11
          61 WLWPNAO-----RDSEERVLVTECGGGDS-IFCKTLTIPRVVGNDTGAYKCSYRDV-- 110
Db
         119 RIEGTTAASSYVFVRDFEQPFINKPDT-----LLVNRKDAMWVPCLVSIPGLNVTL--R 170
Qу
                  Db
         111 ----DIASTVYVYVRDYRSPFIASVSDOHGIVYITENKNKTVVIPCRGSISNLNVSLCAR 166
         171 SQSSVLWPDGQEVVWDDRRGMLVSTPLLHDALYLQCETTWGDQDFLSNPFLVHITGNELY 230
Qу
                   167 YPEKRFVPDGNRISWDSEIGFTLPSYMISYAGMVFCEAKINDETYOSIMYIVVVVGYRIY 226
Db
         231 DIQLLPRKSLELLVGEKLVLNCTVWAEFNSGVTFDWDYPGKQAERGKWVPERRSQQTHTE 290
Qy
             227 DVILSPPHEIELSAGEKLVLNCTARTELNVGLDFTWHSPPSKSHHKKIVNRDVKPFPGTV 286
Db
         291 LS---SILTIHNVSQHDLGSYVCKANNGIQRFRESTEVIVHENPFISVEWLKGPILEATA 347
Qу
                 287 AKMFLSTLTIESVTKSDOGEYTCVASSGRMIKRNRTFVRVHTKPFIAFGSGMKSLVEATV 346
Db
Qy
         348 GDELVKLPVKLAAYPPPEFQWYKDGKALSGRHS---PHALVLKEVTEASTGTYTLALWNS 404
             347 GSQ-VRIPVKYLSYPAPDIKWYRNGRPIESNYTMIVGDELTIMEVTERDAGNYTVILTNP 405
Db
Qу
         405 AAGLRRNISLELVVNVPPQIHEKEASSP-SIYSRHSRQALTCTAYGVPLPLSIQWHWRPW 463
              Db
         406 ISMEKQSHMVSLVVNVPPQIGEKALISPMDSYQYGTMQTLTCTVYANPPLHHIQWYWQLE 465
Qу
         464 TPCKMFAQRSLRRRQQQDLMPQCRDWRAVTTQDAVNPIESLDTWTEFVEGKNKTVSKLVI 523
                              1::|| |
                                          111
                        \perp
                                                       :|||||||||
Db
         466 EACSY-----RPGQTSPYACKEWRHVEDFQGGNKIEVTKNQYALIEGKNKTVSTLVI 517
         524 ONANVSAMYKCVVSNKVGQDERLIYFYVTTIPDGFTIESKPSEELLEGQPVLLSCQADSY 583
Oy
```

Db	518		574
Qу	584	KYEHLRWYRLNLSTLHDAHGNPLLLDCKNVHLFATPLAASLEEVAPGARHATLSLS	639
Db	575	: : : : :: :: TFENLTWYKLGSQATSVHMGESLTPVCKNLDALWKLNGTMFSNSTNDILIVA	626
Qу	640	IPRVAPEHEGHYVCEVQDRRSHDKHCHKKYLSVQALEAPRLTQNLTDLLVNVSDSLEMQC	699
Db	627	::: :::: : : : ::: : FQNASLQDQGDYVCSAQDKKTKKRHCLVKQLIILERMAPMITGNLENQTTTIGETIEVTC	686
Qу	700	LVAGAHAPSIVWYKDERLLEEKSGVDLADSNQKLSIQRVREEDAGRYLCSVCNAKGCVNS	759
Db	687	: : : : : : : : : : : : : : : : : : : : :	746
Qу	760	SASVAVEGSEDKGSMEIVILVGTGVIAVFFWVLLLLIFCNMRRPAHADIKTGYLSIIMDP	819
Db	747	: ::: :: :: : ::: :: ::	806
Qу	820	GEVPLEEQCEYLSYDASQWEFPRERLHLGRVLGYGAFGKVVEASAFGIHKGSSCDTVAVK	879
Db	807	: : : : : : : :	866
Qу	880	MLKEGATASEHRALMSELKILIHIGNHLNVVNLLGACTKPQGPLMVIVEFCKYGNLSNFL	939
Db	867		926
Qy	940	RAKRDAFSPCAEKSPEQRGRFRAMVELARLDRRRPGSSDRVLFARFSKTEGGARRA	995
Db	927	RGKRNEFVPYKSKGARFRQGKDYVGELS-VDLKRRLDSITSSQSSASSGFVEEK	979
QУ	996	SPDQEAEDLWLSPLTMEDLVCYSFQVARGMEFLASRKCIHRDLAARNILLSESDV ::: : : : : : :	1050
Db	980	SLSDVEEEEASEELYKDFLTLEHLICYSFQVAKGMEFLASRKCIHRDLAARNILLSEKNV	1039
Qу	1051	VKICDFGLARDIYKDPDYVRKGSARLPLKWMAPESIFDKVYTTQSDVWSFGVLLWEIFSL	1110
Db	1040	VKICDFGLARDIYKDPDYVRKGDARLPLKWMAPETIFDRVYTIQSDVWSFGVLLWEIFSL	1099
Qу	1111	GASPYPGVQINEEFCQRLRDGTRMRAPELATPAIRRIMLNCWSGDPKARPAFSELVEILG	1170
Db	1100	GASPYPGVKIDEEFCRRLKEGTRMRAPDYTTPEMYQTMLDCWHEDPNQRPSFSELVEHLG	1159
Qy	1171	DLLQGRGLQEEEEVCMAPRS-SQSSEEGSFSQVSTMALHIAQADAEDSPPSLQRHSLAAR : : : : : : :	1229
Db	1160	NLLQANAQQDGKDYIVLPMSETLSMEEDSGLSLPTSPVSCMEEEEVCDPKFH	1211
Qy	1230	YYNWVSFPGCLARGAETRGSSRMKTFEEFPM-TPTTYKGSVDNQTDSGMVLASEEFEQIE	1288
Db	1212	YDNTAGISHYLQNSKRKSRPVSVKTFEDIPLEEPEVKVIPDDSQTDSGMVLASEELKTLE	1271
Qy	1289	SRHRQESGFSCKGPGQNVAVTRAHPDSQGRRRRPERGARGGQVFYNSE 1336 :: : : : :	
Db	1272	DRNKLSPSFGGMMPSKSRESVASEGSNQTSGYQSGYHSD 1310	